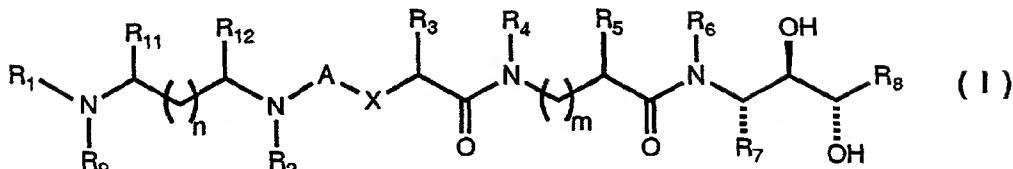


What Is Claimed Is:

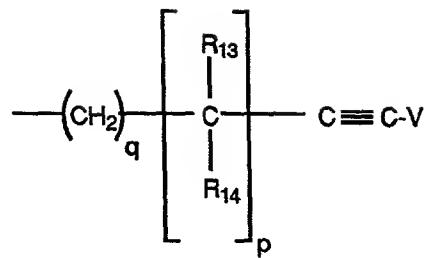
1. A compound of Formula I:



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wherein A is selected from methylene, CO, SO and SO₂;
 wherein X is selected from oxygen atom, methylene and
~~NR₁₀~~ with R₁₀ selected from hydrido, alkyl and benzyl;
 10 wherein each of R₁ and R₉ is a group independently
 selected from hydrido, alkyl, cycloalkyl, alkoxyacetyl,
 haloalkyl, alkoxycarbonyl, benzyloxycarbonyl,
 loweralkanoyl, haloalkylacyl, phenyl, benzyl, naphthyl,
 and naphthylmethyl, any one of which groups having a
 15 substitutable position may be optionally substituted with
 one or more radicals selected from alkyl, alkoxy, alkenyl,
 alkynyl, halo, haloalkyl, cyano and phenyl, and wherein
 the nitrogen atom to which R₁ and R₉ are attached may be
 combined with oxygen to form an N-oxide; wherein R₂ is
 20 selected from hydrido, alkyl, dialkylaminoalkyl,
 alkylacylaminoalkyl, benzyl and cycloalkyl; wherein R₃ is
 selected from alkyl, cycloalkylalkyl, acylaminoalkyl,
 phenylalkyl, naphthylmethyl, aryl, heterocyclicalkyl and
 heterocycliccycloalkyl, wherein the cyclic portion of any
 25 of said phenylalkyl, naphthylmethyl, aryl,
 heterocyclicalkyl and heterocycliccycloalkyl groups may be
 substituted by one or more radicals selected from halo,
 hydroxy, alkoxy and alkyl; wherein each of R₄ and R₆ is
 independently selected from hydrido, alkyl, benzyl and
 30 cycloalkyl; wherein each of R₅ and R₈ is independently
 selected from

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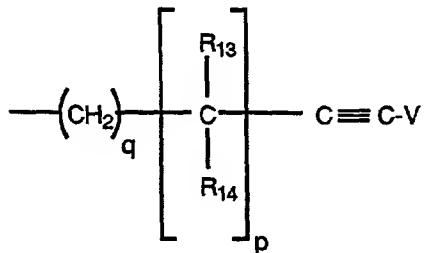
wherein V is selected from hydrido, alkyl, cycloalkyl, haloalkyl, benzyl and phenyl; wherein each of R₁₃ and

5 R₁₄ is a radical independently selected from hydrido, alkyl, alkenyl, alkynyl, cycloalkyl, phenyl, heterocyclic, heterocyclicalkyl and heterocycliccycloalkyl; wherein R₇ is selected from substituted or unsubstituted alkyl, cycloalkyl, phenyl, 10 cycloalkylalkyl and phenylalkyl, any one of which may be substituted with one or more groups selected from alkyl, hydroxy, alkoxy, halo, haloalkyl, alkenyl, alkynyl and cyano; wherein each of R₁₁ and R₁₂ is independently selected from hydrido, alkyl, haloalkyl, dialkylamino 15 and phenyl; and wherein m is zero or one; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

20

2. Compound of Claim 1 wherein A is selected from methylene, CO, SO and SO₂; wherein X is selected from oxygen atom, methylene and NR_{10} with R₁₀ selected from hydrido, alkyl and benzyl; wherein each of R₁ and R₉ is 25 independently selected from hydrido, lower alkyl, haloalkyl, cycloalkyl, alkoxycarbonyl, benzyloxycarbonyl, loweralkanoyl, alkoxyacetyl, phenyl and benzyl, and wherein the nitrogen atom to which R₁ and R₉ are attached may be combined with oxygen to form an N-oxide; wherein each of R₂, R₄ and R₆ is independently selected from hydrido and 30 alkyl; wherein R₃ is selected from phenylalkyl, naphthylmethyl, cyclohexylalkyl, cyclopentylalkyl,

heteroarylalkyl and heteroarylcycloalkyl; wherein each of R₅ and R₈ is independently selected from



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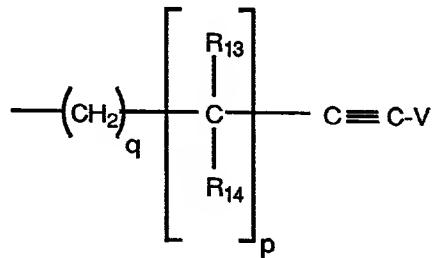
wherein V is selected from hydrido, alkyl, haloalkyl, benzyl and phenyl; wherein each of R₁₃ and R₁₄ is a radical independently selected from hydrido, alkyl, alkenyl, alkynyl, cycloalkyl, heteroaryl,

- 10 heteroarylalkyl and heteroarylcycloalkyl; wherein R₇ is selected from substituted or unsubstituted cyclohexylmethyl and benzyl, either one of which may be substituted with one or more groups selected from alkyl, hydroxy, alkoxy, halo and haloalkyl; wherein each of R₁₁ and R₁₂ is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero or one; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through
- 20 five; or a pharmaceutically-acceptable salt thereof.

3. Compound of Claim 2 wherein A is selected from methylene, CO, SO and SO₂; wherein X is selected from oxygen atom, methylene and >NR₁₀ with R₁₀ selected from

- 25 hydrido, alkyl and benzyl; wherein each of R₁ and R₉ is independently selected from hydrido, alkyl, alkoxyacyl, haloalkyl, alkoxycarbonyl, benzyloxycarbonyl and benzyl, and wherein the nitrogen atom to which R₁ and R₉ are attached may be combined with oxygen to form an N-oxide;
- 30 wherein each of R₂, R₄ and R₆ is independently selected from hydrido and alkyl; wherein R₃ is selected from benzyl, phenethyl, cyclohexylmethyl, phenpropyl, pyrrolidinyl, piperidinyl, pyrrolidinylmethyl,

piperidinylmethyl, pyrazolemethyl, pyrazoleethyl,
 pyridylmethyl, pyridylethyl, thiazolemethyl,
 thiazoleethyl, imidazolemethyl, imidazoleethyl,
 thienylmethyl, thienylethyl, furanylmethyl, furanylethyl,
 5 oxazolemethyl, oxazoleethyl, isoxazolemethyl,
 isoxazoleethyl, pyridazinemethyl, pyridazineethyl,
 pyrazinemethyl and pyrazineethyl; wherein each of R₅ and
 R₈ is independently selected from



wherein V is selected from hydrido, alkyl and haloalkyl;
 wherein each of R₁₃ and R₁₄ is a radical independently
 selected from hydrido, alkyl, alkenyl, alkynyl, thiazole
 and thiazolemethyl; wherein R₇ is cyclohexylmethyl;
 wherein each of R₁₁ and R₁₂ is independently selected
 from hydrido, alkyl, dialkylamino and phenyl; wherein m
 is zero or one; wherein n is a number selected from zero
 through five; wherein p is a number selected from zero
 15 through five; and wherein q is a number selected from
 zero through five; or a pharmaceutically-acceptable salt
 thereof.

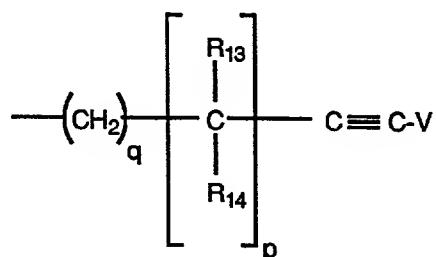
4. Compound of Claim 3 wherein A is selected
 25 from CO and SO₂; wherein X is selected from oxygen atom,
 methylene and >NR₁₀ with R₁₀ selected from hydrido and
 methyl; wherein each of R₁ and R₉ is independently
 selected from hydrido, lower alkyl, alkoxyacyl,
 alkoxy carbonyl, benzyloxycarbonyl, haloalkyl and benzyl,
 30 and wherein the nitrogen atom to which R₁ and R₉ are
 attached may be combined with oxygen to form an N-oxide;
 wherein R₂ is selected from hydrido, methyl, ethyl and
 isopropyl; wherein R₃ is selected from benzyl, phenethyl,

cyclohexylmethyl, pyrrolidinyl, piperidinyl,
 pyrrolidinylmethyl, piperidinylmethyl, pyrazolemethyl,
 pyrazoleethyl, pyridylmethyl, pyridylethyl,

thiazolemethyl, thiazoleethyl, imidazolemethyl,

5 imidazoleethyl, thienylmethyl, thienylethyl,
 furanylmethyl, furanylethyl, oxazolemethyl, oxazoleethyl,
 isoxazolemethyl, isoxazoleethyl, pyridazinemethyl,
 pyridazineethyl, pyrazinemethyl and pyrazineethyl;

10 wherein each of R₄ and R₆ is independently selected from
 hydrido and methyl; wherein each of R₅ and R₈ is
 independently selected from



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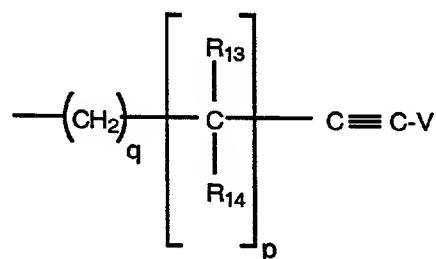
wherein V is selected from hydrido, alkyl and
 trifluoromethyl; wherein each of R₁₃ and R₁₄ is a
 radical independently selected from hydrido, alkyl and
 alkynyl; wherein R₇ is cyclohexylmethyl; wherein each of

20 R₁₁ and R₁₂ is independently selected from hydrido,
 alkyl, dialkylamino and phenyl; wherein m is zero;
 wherein n is a number selected from zero through five;
 wherein p is a number selected from zero through five;
 and wherein q is a number selected from zero through
 25 five; or a pharmaceutically-acceptable salt thereof.

30 5. Compound of Claim 4 wherein A is selected
 from CO and SO₂; wherein X is selected from oxygen atom
 and methylene; wherein each of R₁ and R₉ is independently
 selected from hydrido, methyl, ethyl, n-propyl, isopropyl,
 benzyl, b, b, b-trifluoroethyl, t-butyloxycarbonyl and
 methoxymethylcarbonyl, and wherein the nitrogen atom to
 which R₁ and R₉ are attached may be combined with oxygen

to form an N-oxide; wherein R₂ is selected from hydrido, methyl, ethyl and isopropyl; wherein R₃ is selected from benzyl, cyclohexylmethyl, phenethyl, pyrazolemethyl, pyrazoleethyl, pyridylmethyl, pyridylethyl,

5 thiazolemethyl, thiazoleethyl, imidazolemethyl, imidazoleethyl, thiethylmethyl, thiethyl, furanylmethyl, furylethyl, oxazolemethyl, oxazoleethyl, isoxazolemethyl, isoxazoleethyl, pyridazinemethyl, pyridazineethyl, pyrazinemethyl and pyrazineethyl; wherein
 10 each of R₅ and R₈ is independently selected from

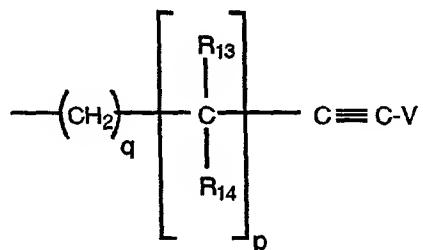


wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R₁₃ and R₁₄ is a radical independently selected from hydrido, methyl, ethyl, propyl and ethynyl; wherein R₇ is cyclohexylmethyl; wherein each of R₄ and R₆ is independently selected from hydrido and methyl; wherein
 15 each of R₁₁ and R₁₂ is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero
 20 each of R₁₁ and R₁₂ is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero
 25 each of R₁₁ and R₁₂ is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero
 thereof.

6. Compound of Claim 5 wherein A is selected from CO and SO₂; wherein X is selected from oxygen atom and methylene; wherein each of R₁ and R₉ is a group independently selected from hydrido, methyl, ethyl, n-propyl, isopropyl, benzyl, b, b, b-trifluoroethyl, t-butyloxycarbonyl and methoxymethylcarbonyl, and wherein

the nitrogen atom to which R₁ and R₉ are attached may be combined with oxygen to form an N-oxide; wherein R₂ is selected from hydrido, methyl, ethyl and isopropyl; wherein R₃ is selected from benzyl, cyclohexylmethyl,

5 phenethyl, imidazolemethyl, pyridylmethyl and 2-pyridylethyl; wherein each of R₅ and R₈ is independently selected from

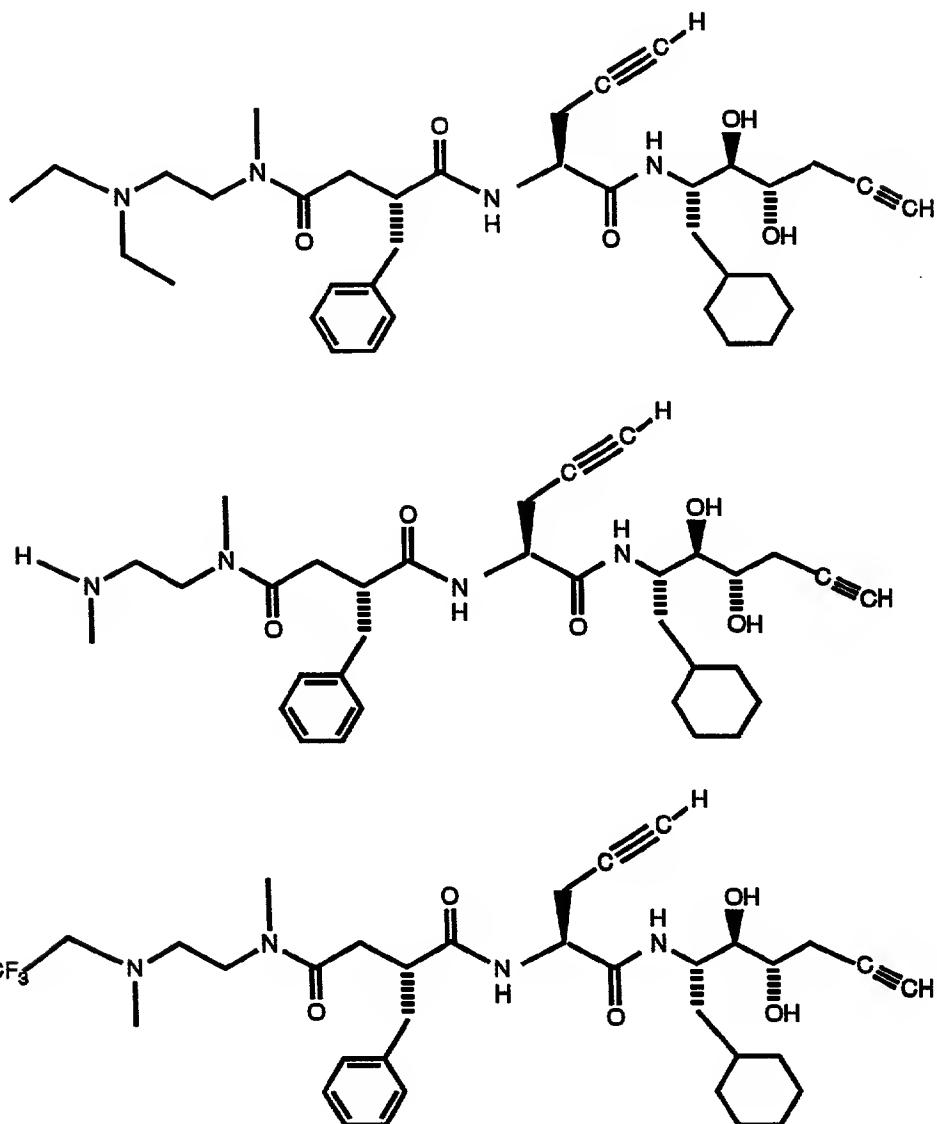


10 wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R₁₃ and R₁₄ is a radical independently selected from hydrido, methyl and ethynyl; wherein R₇ is cyclohexylmethyl; wherein each of R₄ and R₆ is independently selected from hydrido and methyl; wherein each of R₁₁ and R₁₂ is independently selected from hydrido, alkyl and phenyl; wherein m is zero; wherein n is a number selected from zero through three; wherein p is a number selected from one through three; and wherein q is 15 zero or one; or a pharmaceutically-acceptable salt thereof.

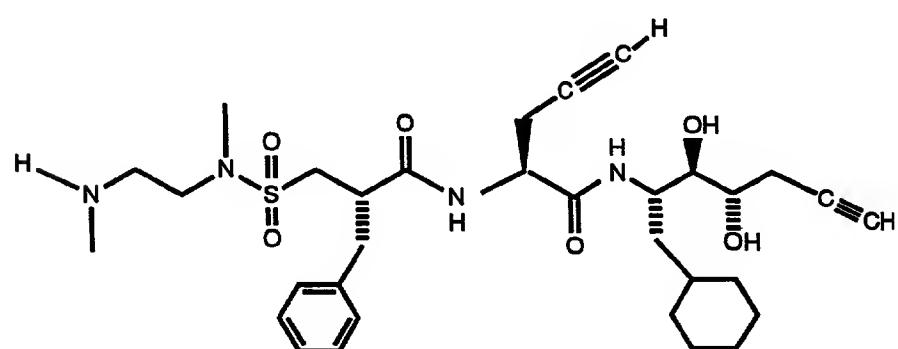
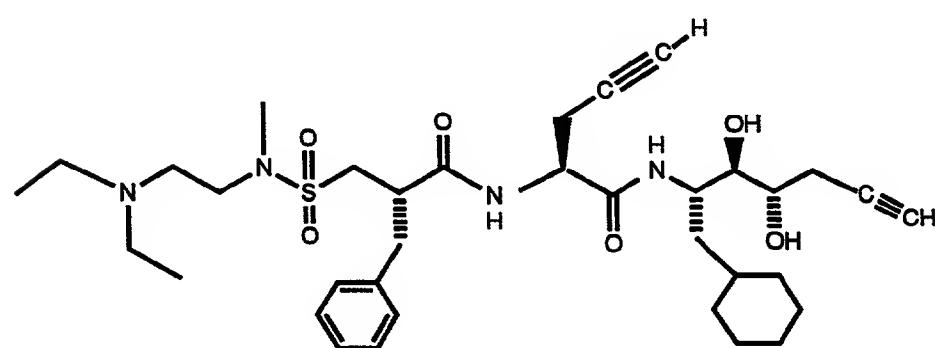
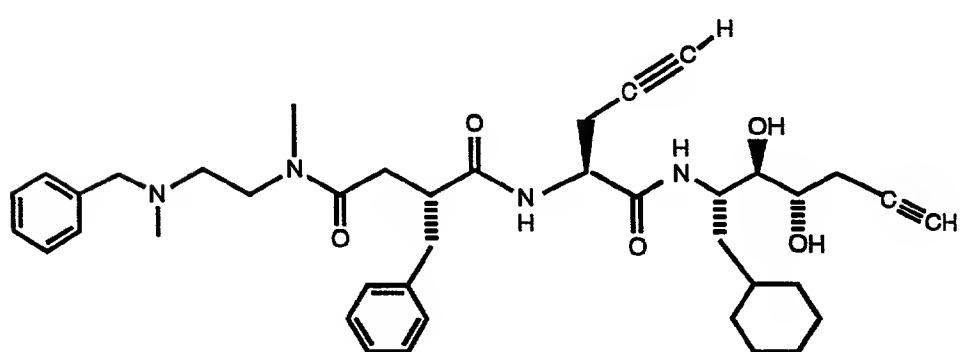
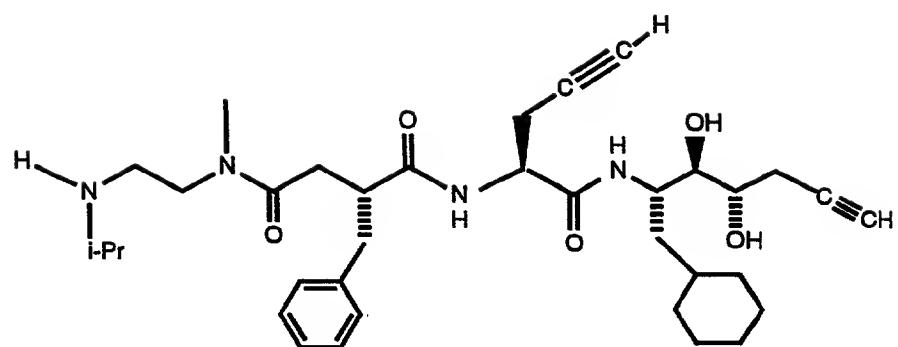
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7. Compound of Claim 6 selected from compounds, their tautomers, and the pharmaceutically-acceptable esters and salts thereof, of the group consisting of

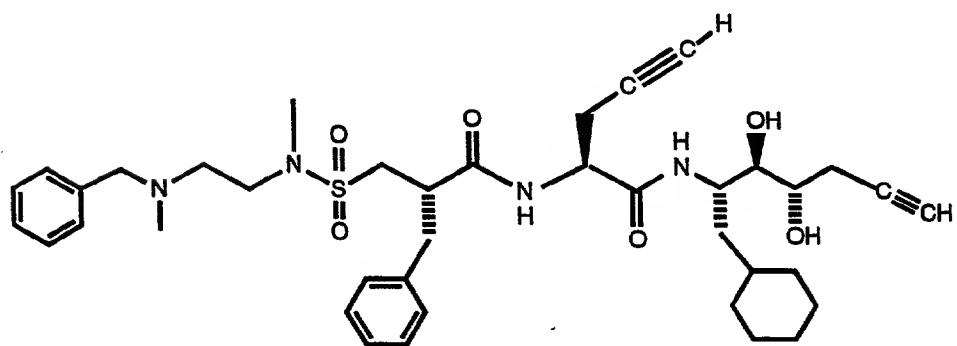
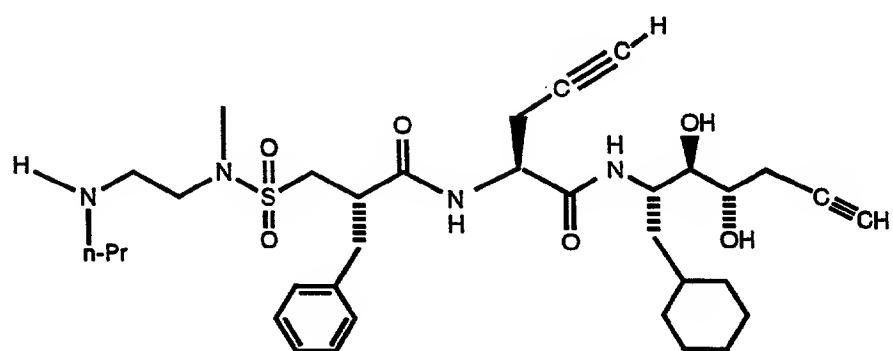
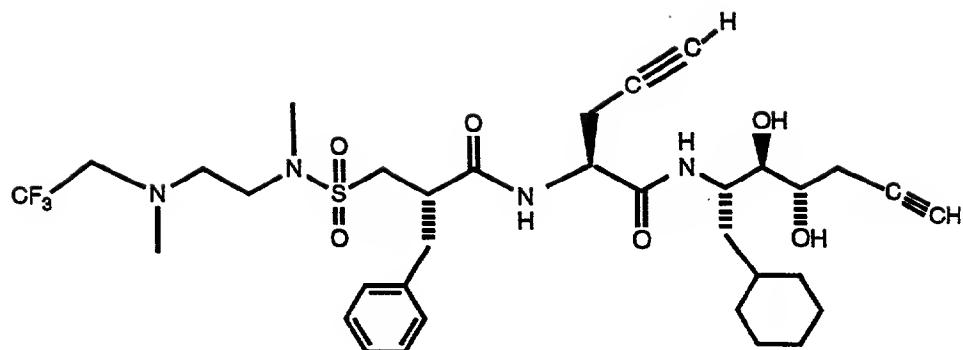
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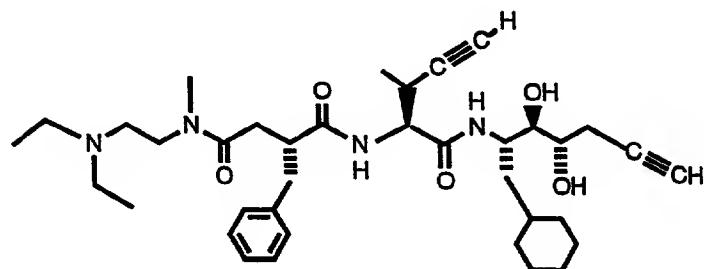
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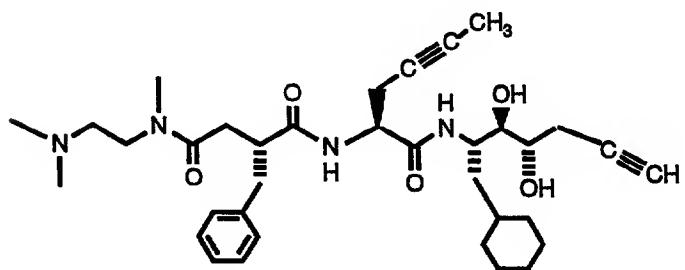
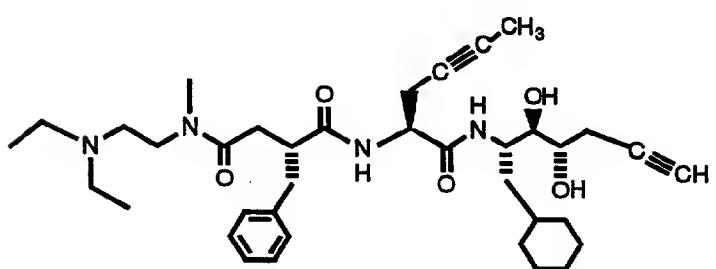
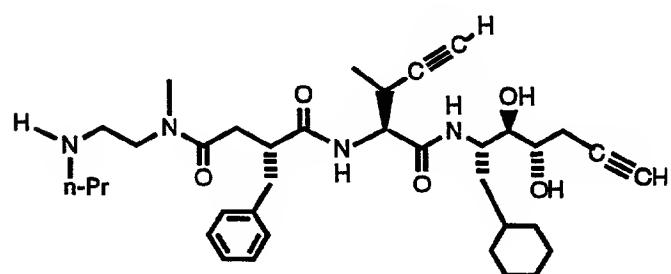
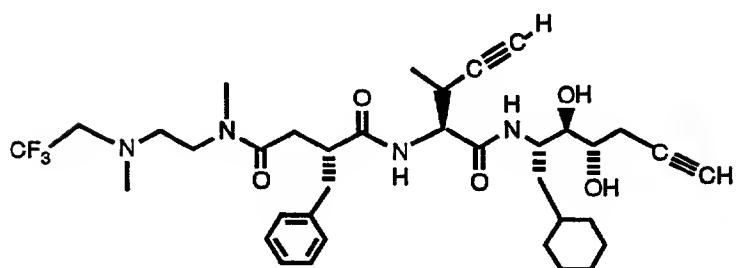
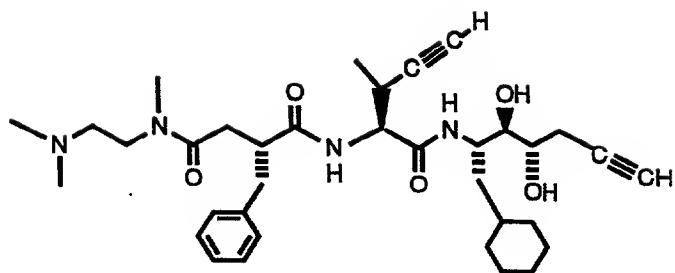


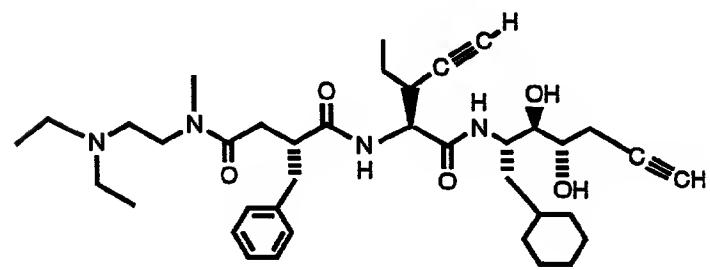
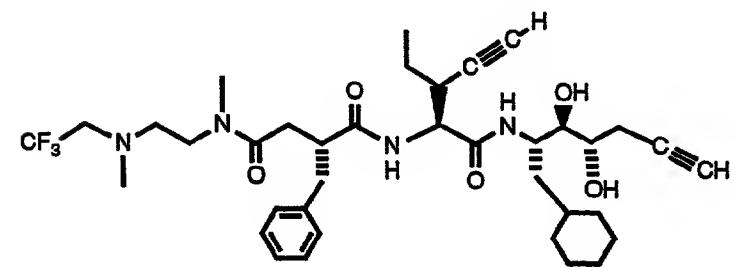
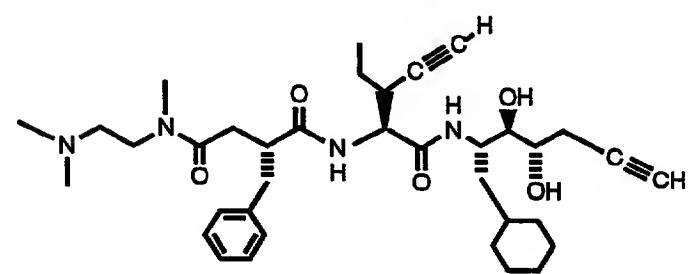
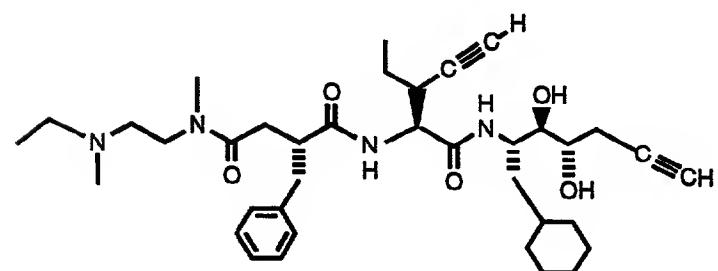
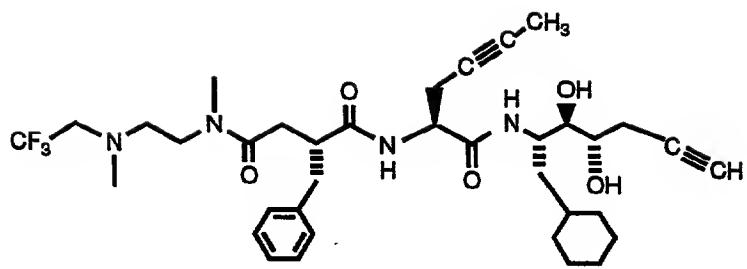
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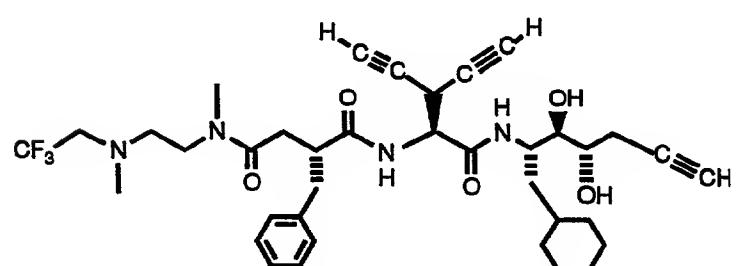
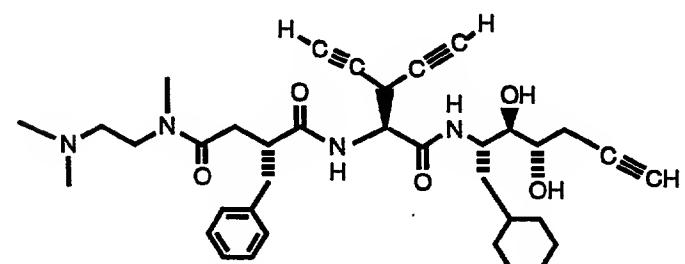
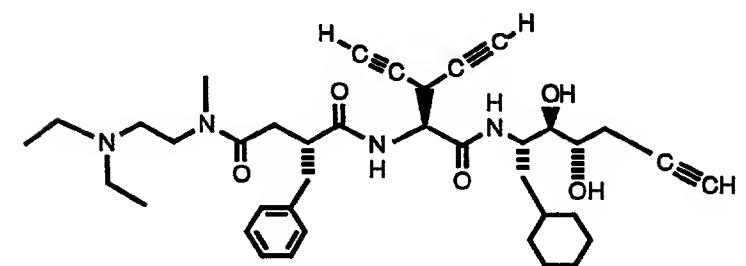
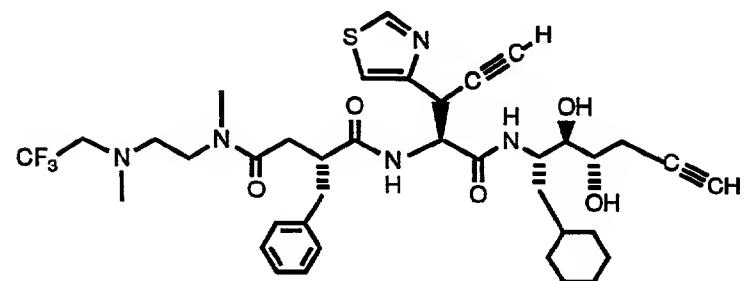
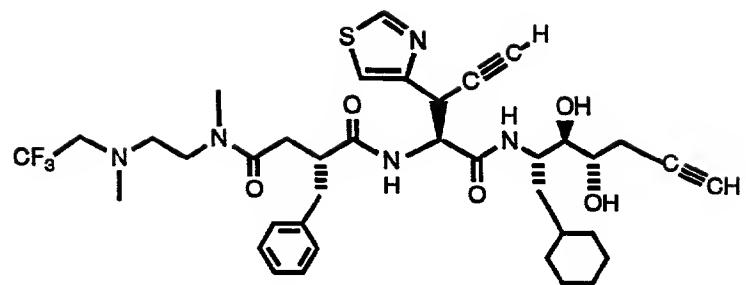




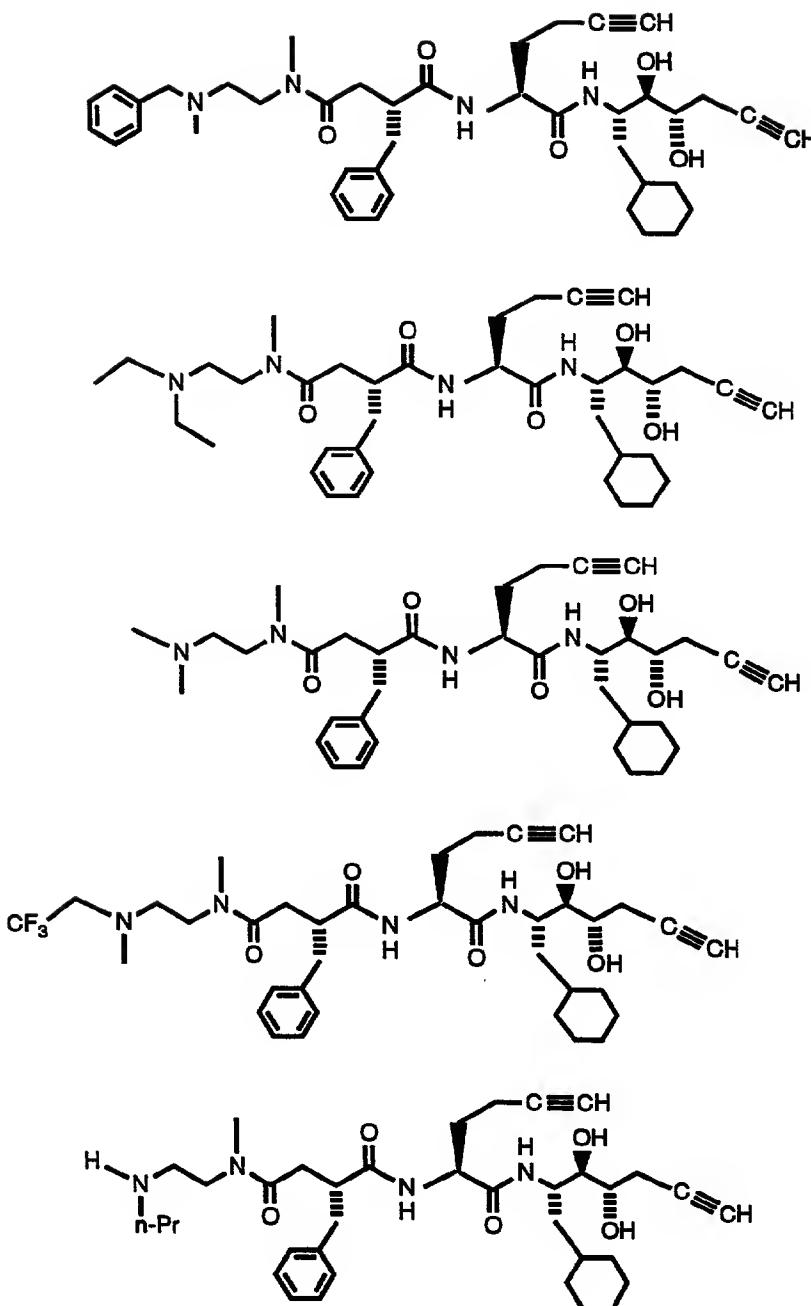
STRUCTURE 5

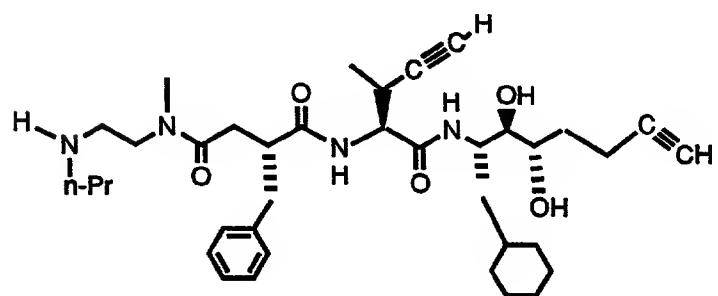
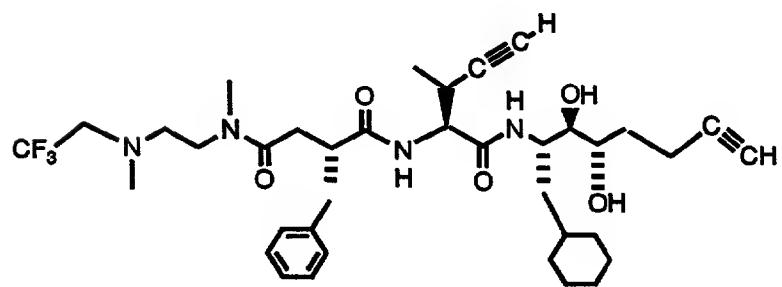
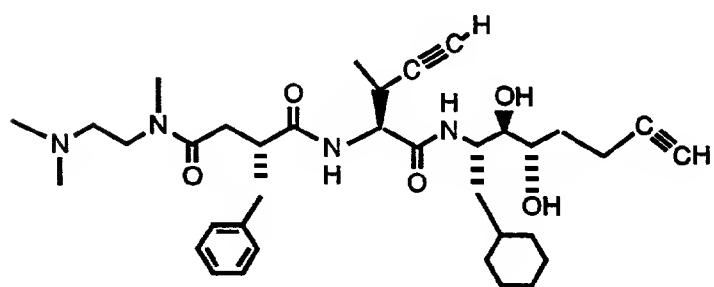
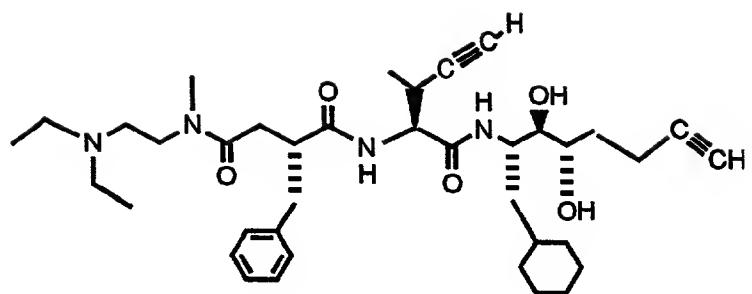
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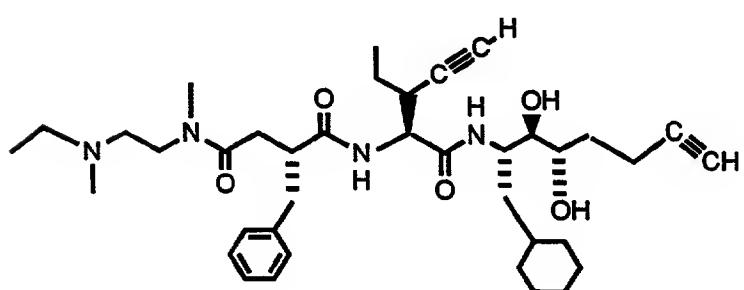
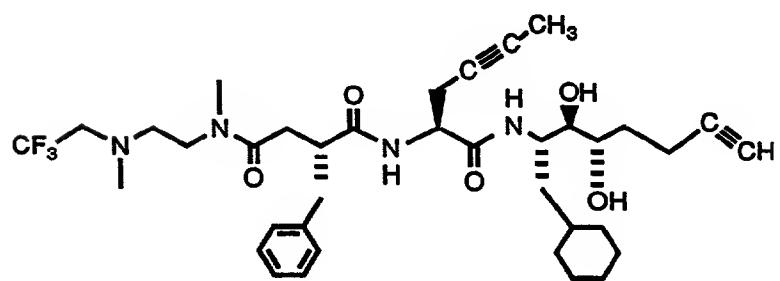
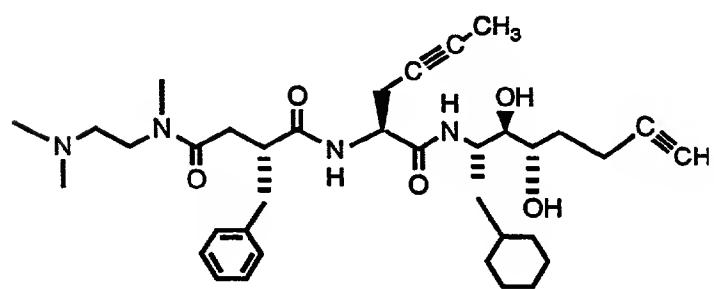
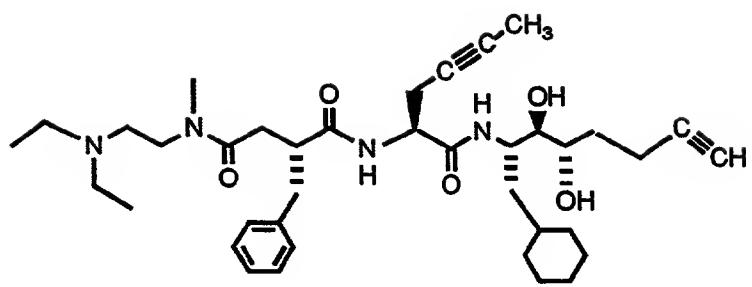
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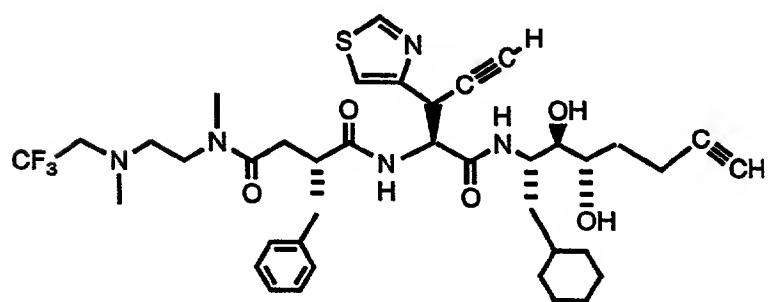
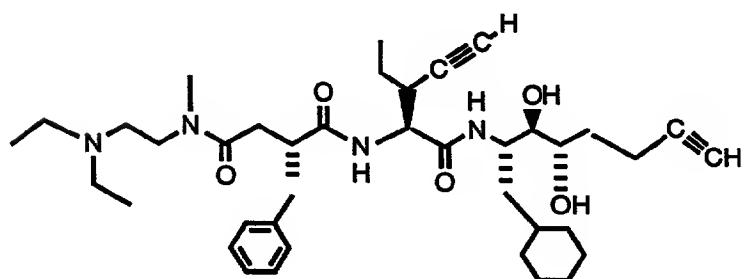
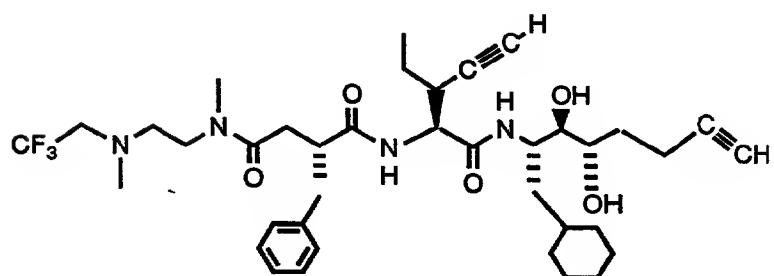
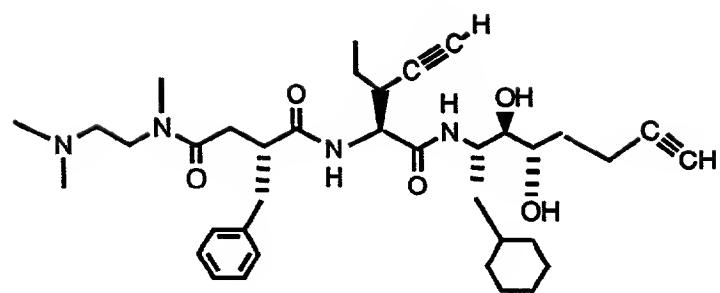


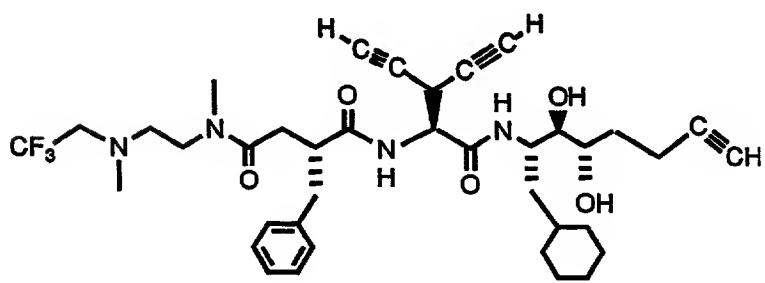
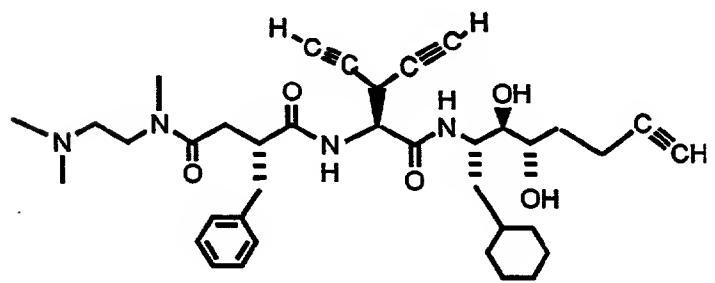
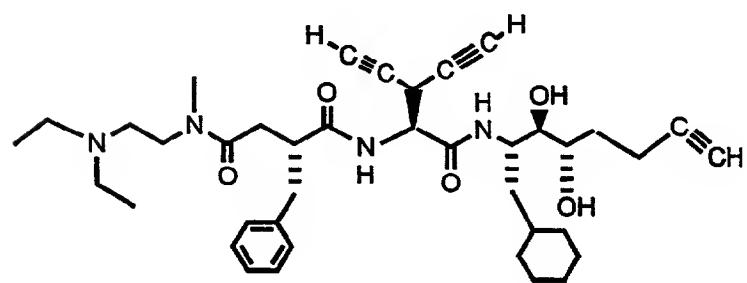
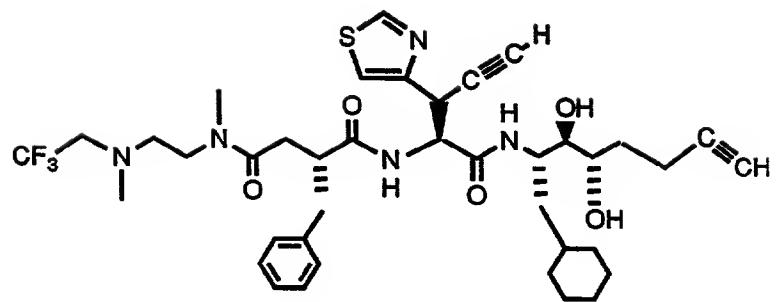
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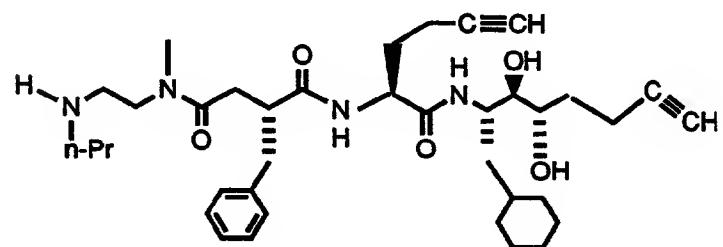
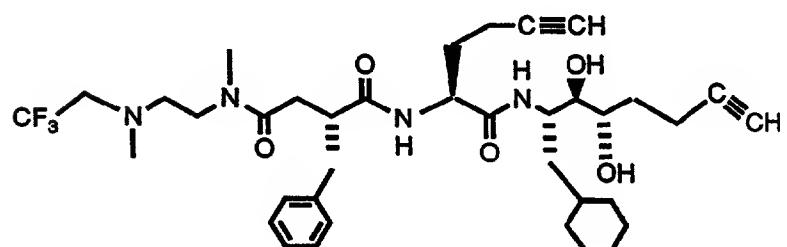
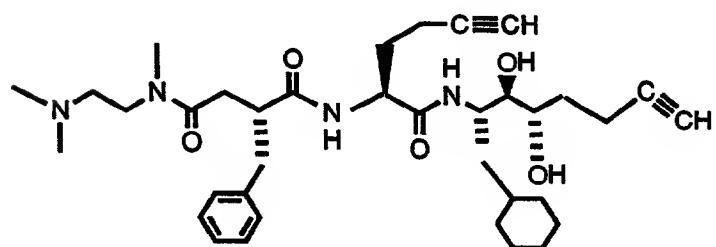
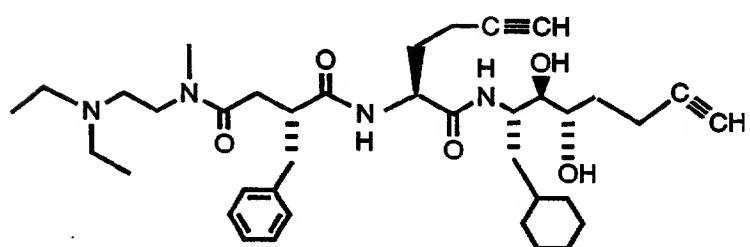
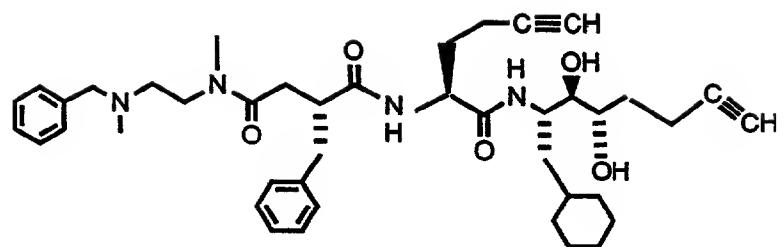


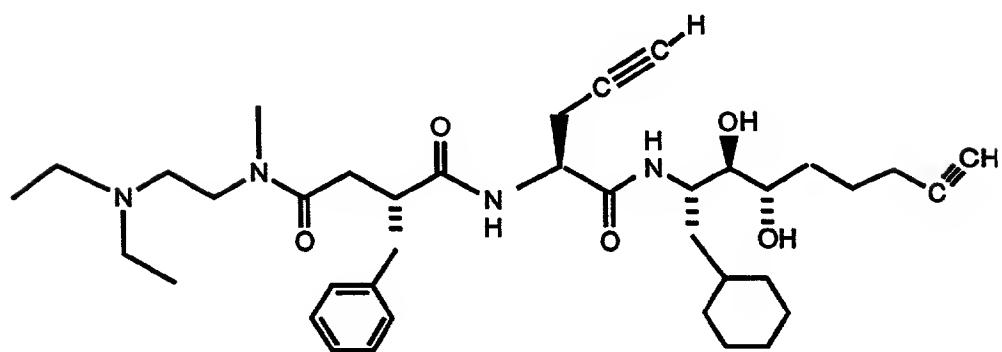
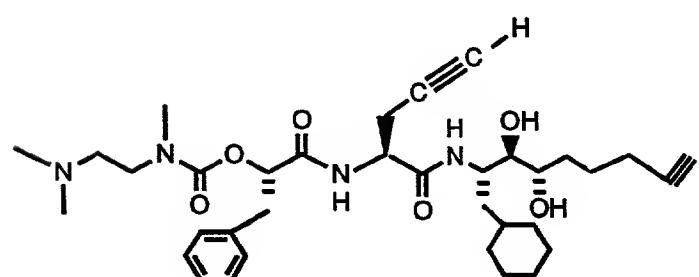
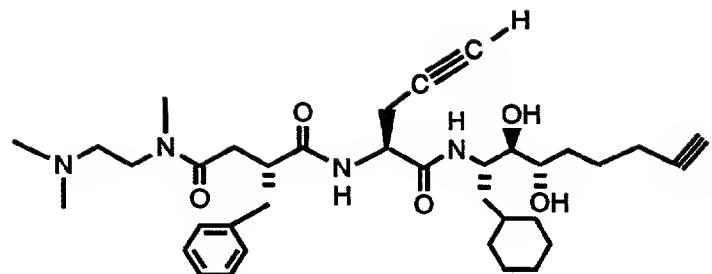




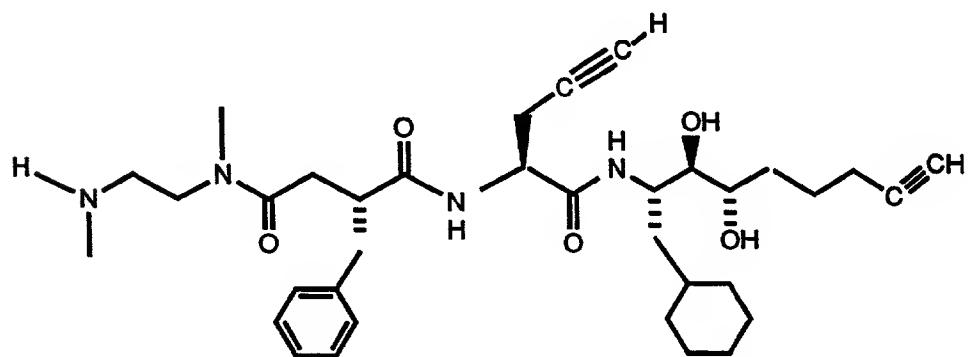


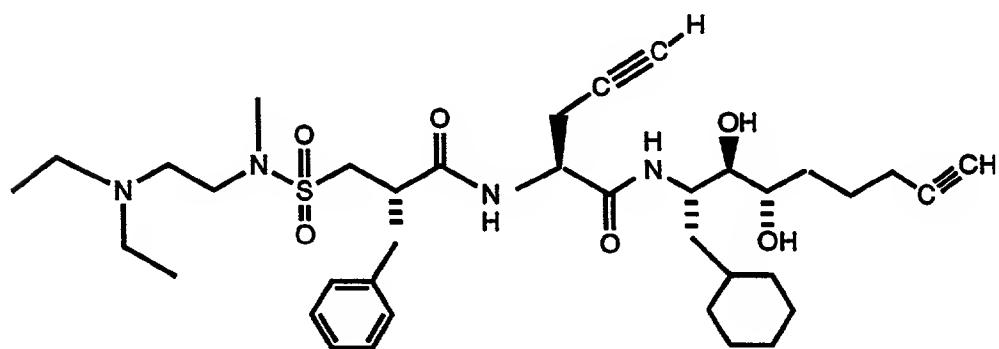
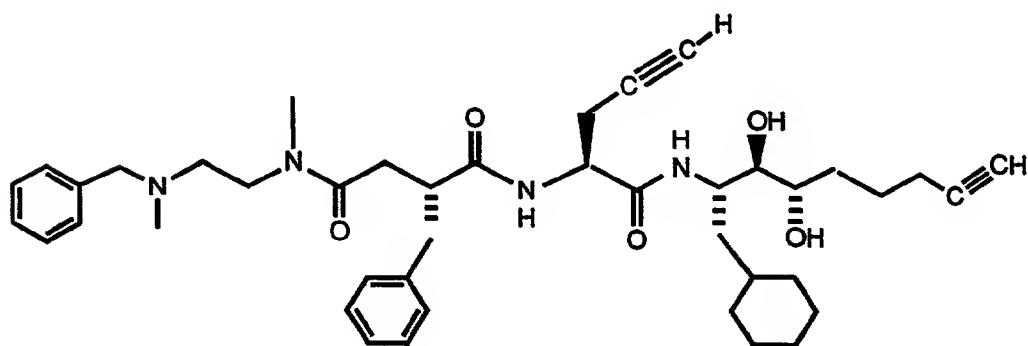
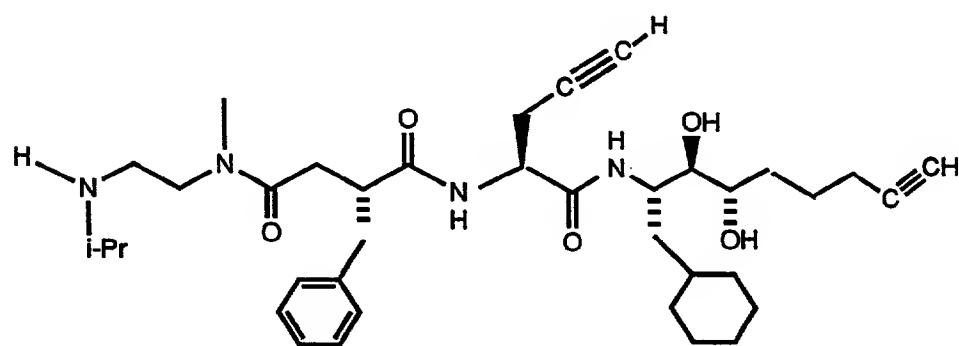
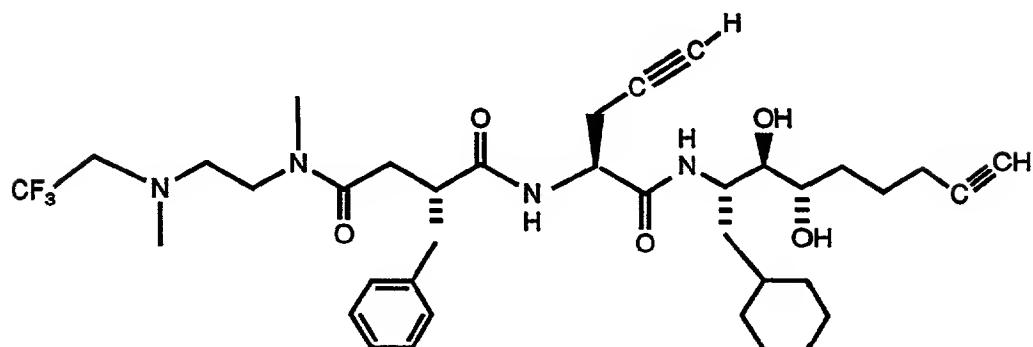




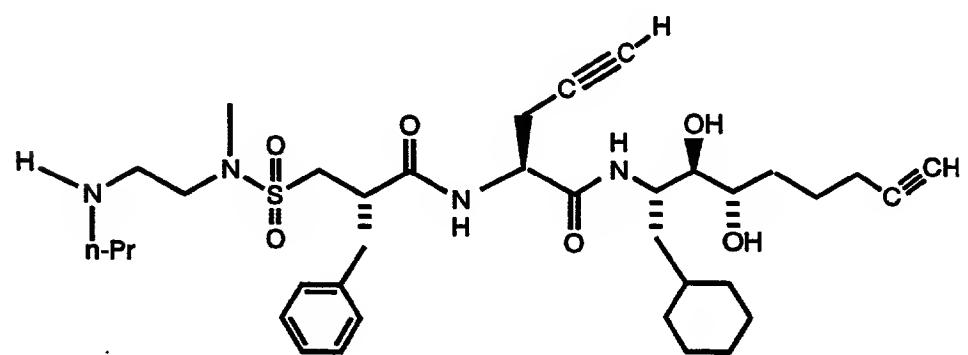
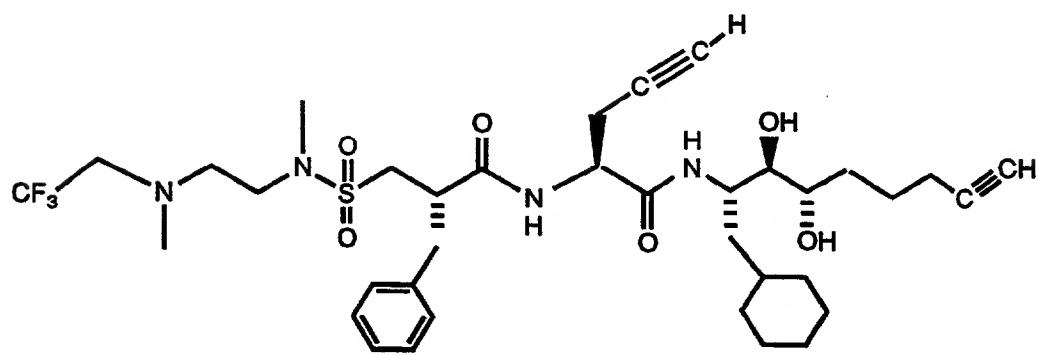
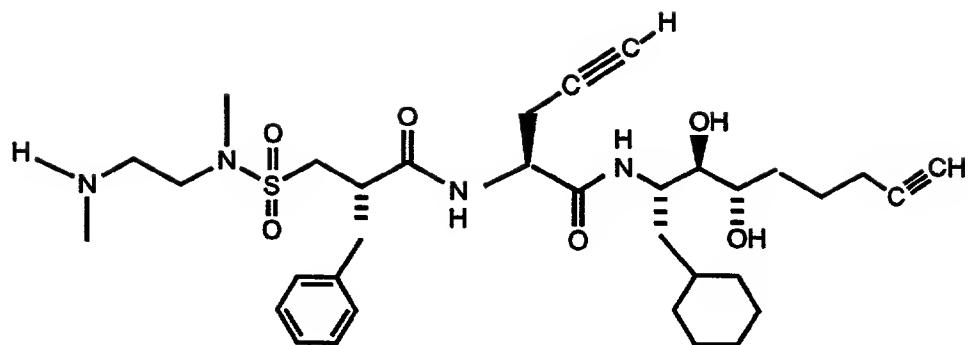


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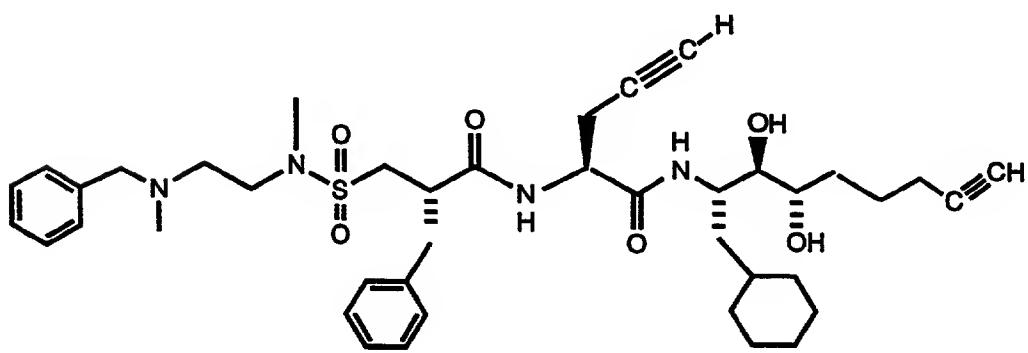


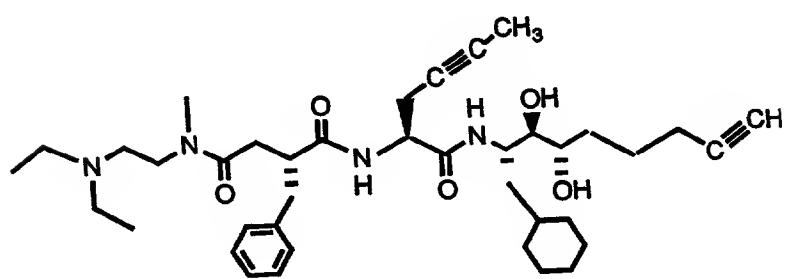
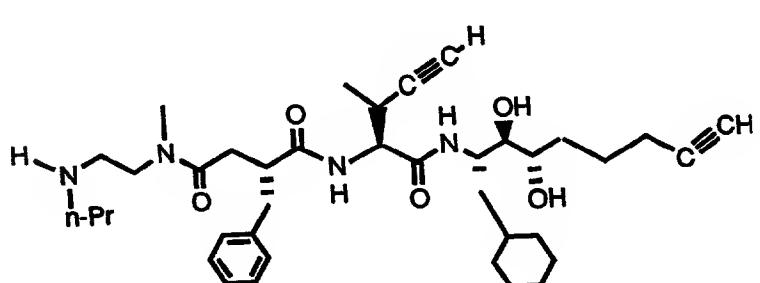
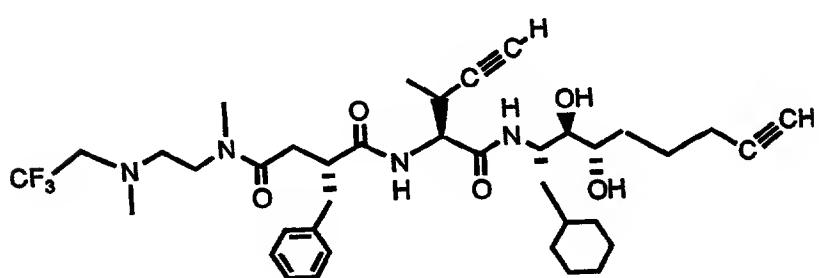
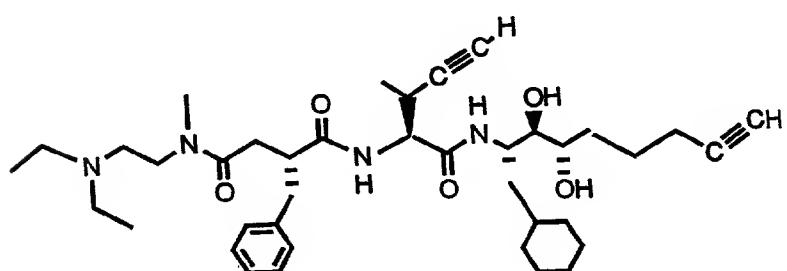
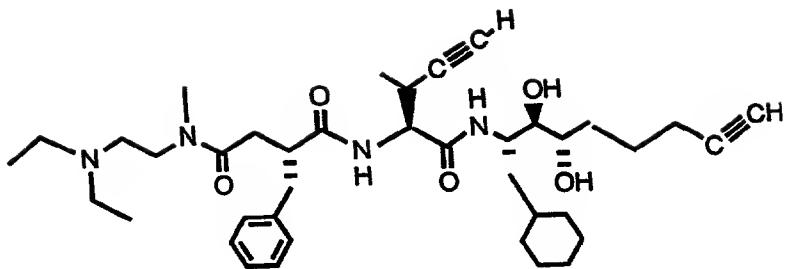


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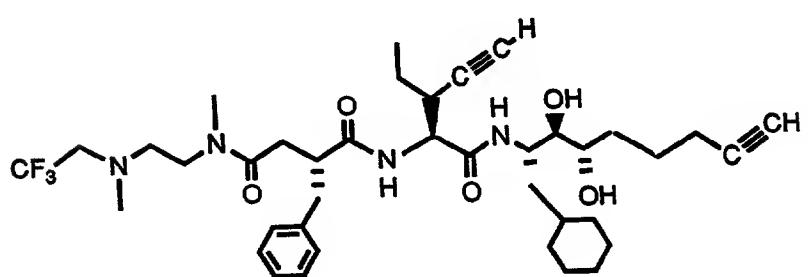
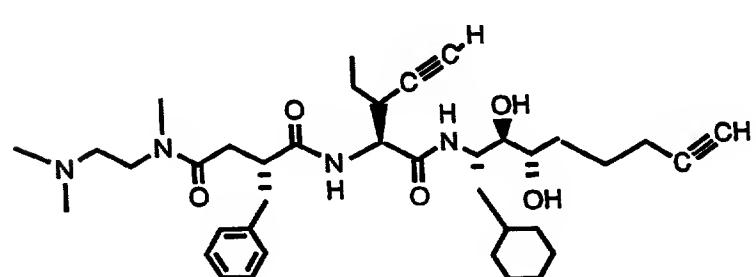
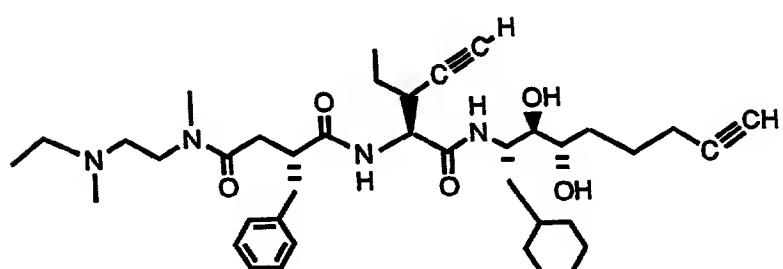
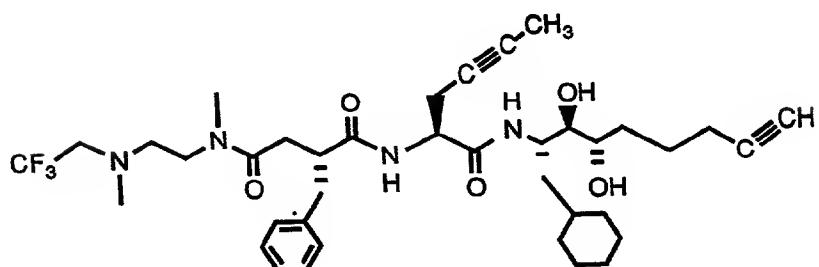
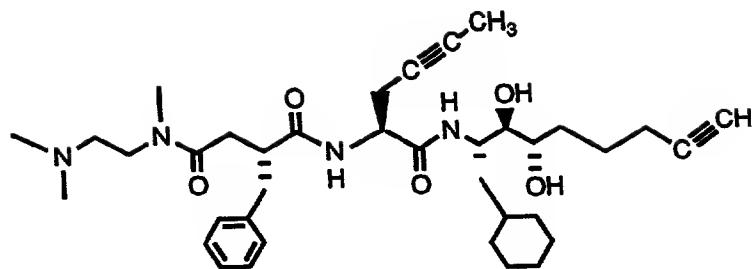


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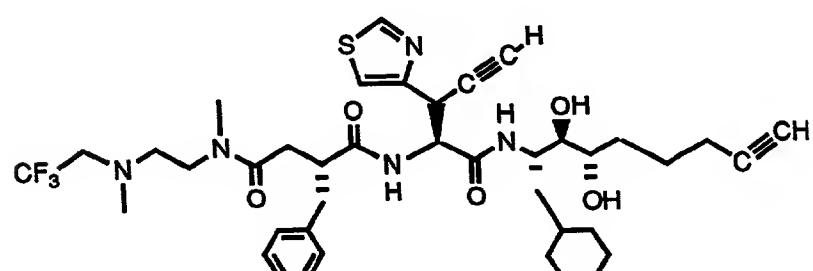
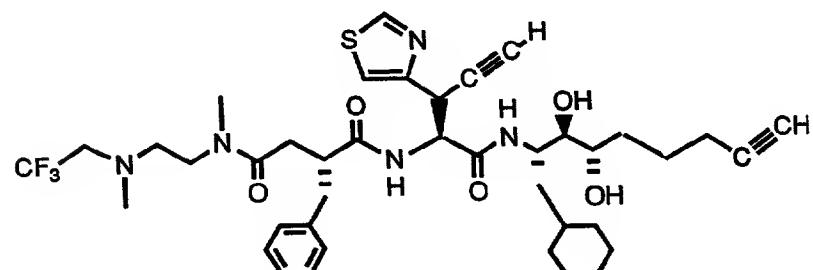
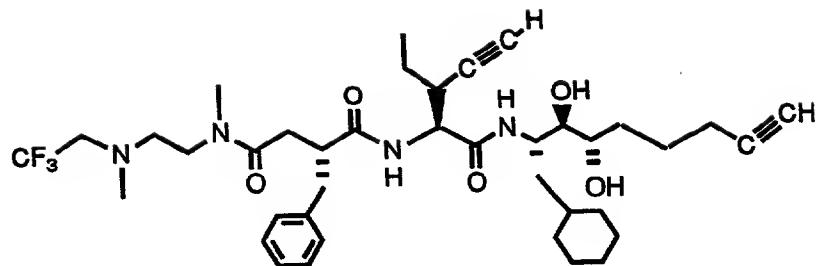




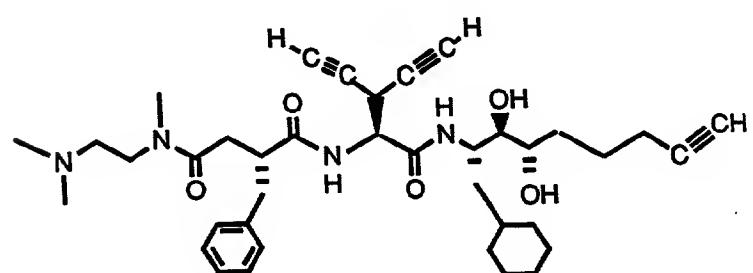
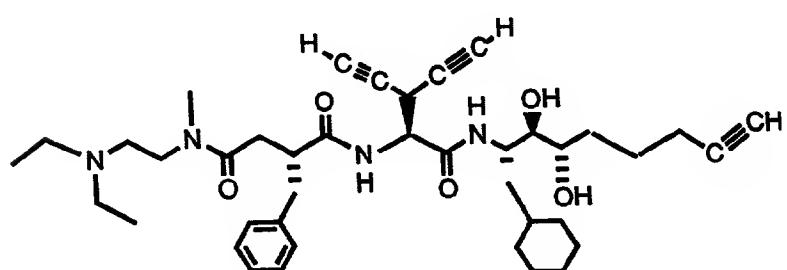
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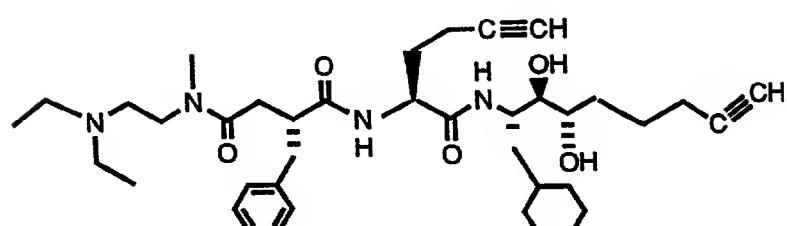
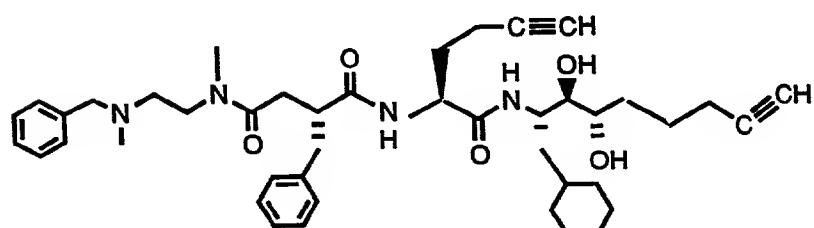
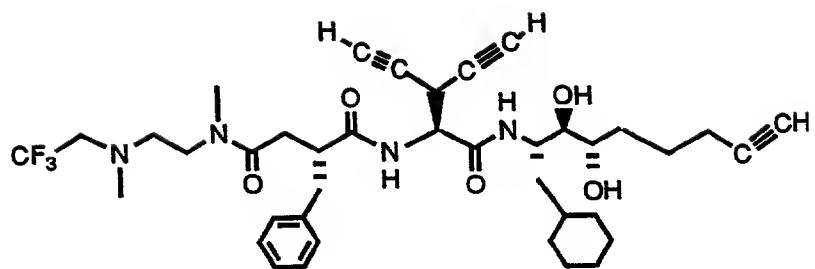


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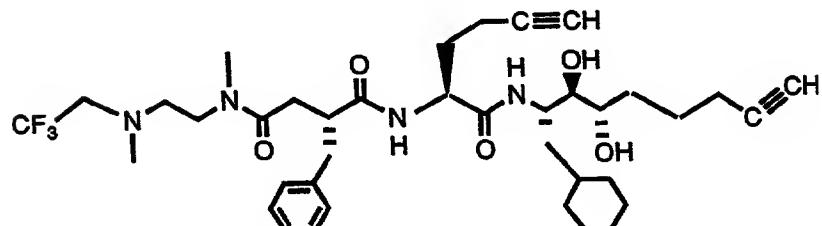
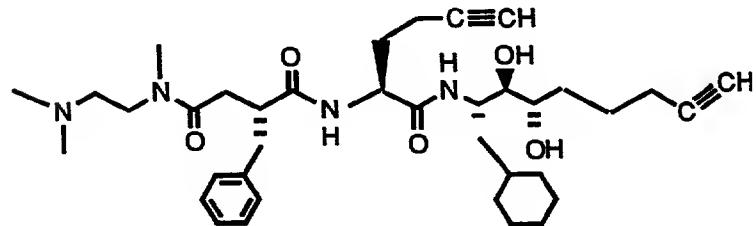


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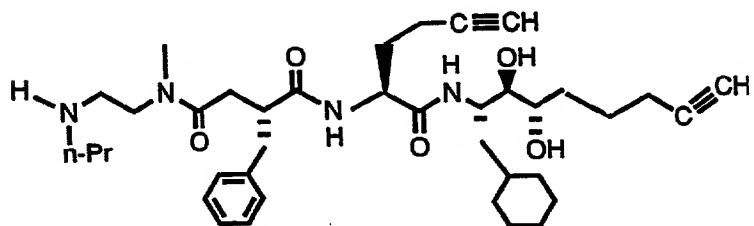
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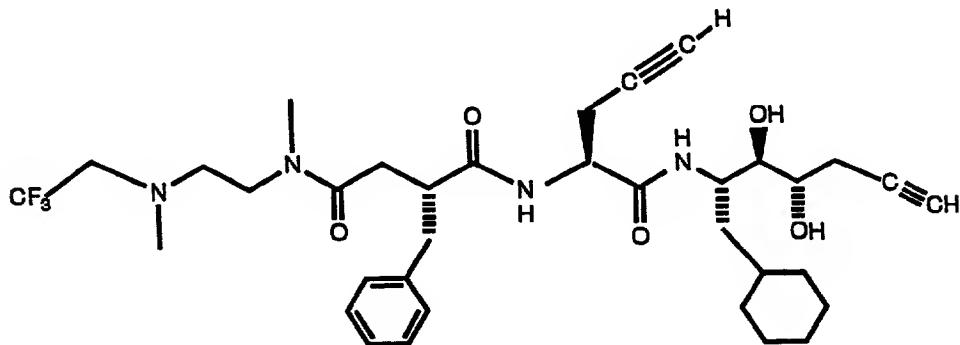
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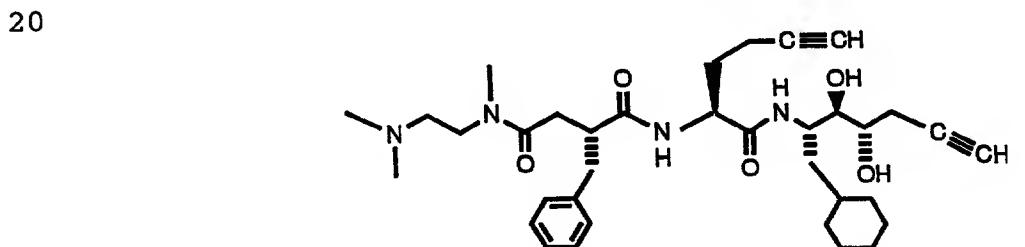
8. Compound of Claim 6 which is N1-[1R*-
 [[[1S,1R*-(cyclohexylmethyl)-2S*,3R*-dihydroxy-
 hexynyl]amino]carbonyl]-3-butynyl]-N4-[2-
 5 (dimethylamino)ethyl]-N4-methyl-2S*-
 (phenylmethyl)butanediamide or a pharmaceutically-
 acceptable salt thereof.

9. Compound of Claim 6 which is [1R*-
 10 [[[1R*-[[[[1S,1R*-(cyclohexylmethyl)-2S*,3R*-dihydroxy-
 hexynyl]amino]carbonyl]-3-butynyl]amino]carbonyl]-2-
 phenylethyl][2-(dimethylamino)ethyl]methylcarbamate or a
 pharmaceutically-acceptable salt thereof.

15 10. Compound of Claim 6 which is

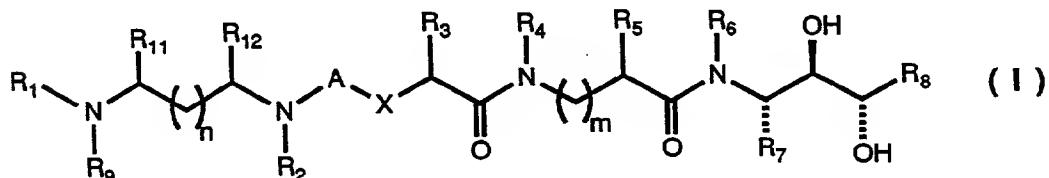


11. Compound of Claim 6 which is



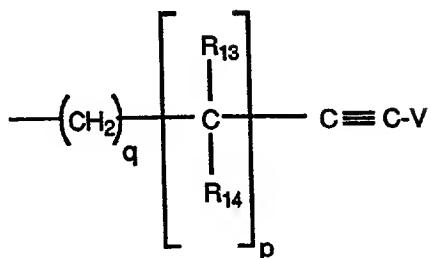
or a pharmaceutically-acceptable salt thereof.

12. A pharmaceutical composition comprising
 a therapeutically-effective amount of a renin-inhibiting
 compound and a pharmaceutically-acceptable carrier or
 diluent, said renin-inhibiting compound selected from a
 5 family of compounds of Formula I:



wherein A is selected from methylene, CO, SO and SO₂;
 10 wherein X is selected from oxygen atom, methylene and
~~>NR₁₀~~ with R₁₀ selected from hydrido, alkyl and benzyl;
 wherein each of R₁ and R₉ is a group independently
 selected from hydrido, alkyl, cycloalkyl, alkoxyacetyl,
 haloalkyl, alkoxy carbonyl, benzyloxycarbonyl,
 loweralkanoyl, haloalkylacyl, phenyl, benzyl, naphthyl,
 and naphthylmethyl, any one of which groups having a
 15 substitutable position may be optionally substituted
 with one or more radicals selected from alkyl, alkoxy,
 alkenyl, alkynyl, halo, haloalkyl, cyano and phenyl, and
 wherein the nitrogen atom to which R₁ and R₉ are
 20 attached may be combined with oxygen to form an N-oxide;
 wherein R₂ is selected from hydrido, alkyl,
 dialkylaminoalkyl, alkylacylaminoalkyl, benzyl and
 cycloalkyl; wherein R₃ is selected from alkyl,
 25 cycloalkylalkyl, acylaminoalkyl, phenylalkyl,
 naphthylmethyl, aryl, heterocyclicalkyl and
 heterocycliccycloalkyl, wherein the cyclic portion of
 any of said phenylalkyl, naphthylmethyl, aryl,
 heterocyclicalkyl and heterocycliccycloalkyl groups may
 30 be substituted by one or more radicals selected from
 halo, hydroxy, alkoxy and alkyl; wherein each of R₄ and
 R₆ is independently selected from hydrido, alkyl, benzyl

and cycloalkyl; wherein each of R₅ and R₈ is independently selected from



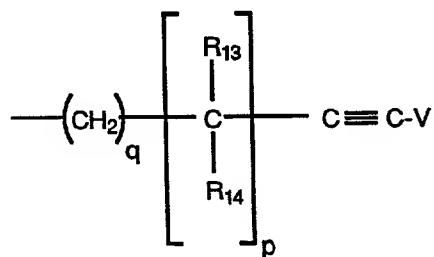
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wherein V is selected from hydrido, alkyl, cycloalkyl, haloalkyl, benzyl and phenyl; wherein each of R₁₃ and R₁₄ is a radical independently selected from hydrido, alkyl, alkenyl, alkynyl, cycloalkyl, phenyl, heterocyclic, heterocyclicalkyl and heterocycliccycloalkyl; wherein R₇ is selected from substituted or unsubstituted alkyl, cycloalkyl, phenyl, cycloalkylalkyl and phenylalkyl, any one of which may be substituted with one or more groups selected from alkyl, hydroxy, alkoxy, halo, haloalkyl, alkenyl, alkynyl and cyano; wherein each of R₁₁ and R₁₂ is independently selected from hydrido, alkyl, haloalkyl, dialkylamino and phenyl; and wherein m is zero or one; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

13. The composition of Claim 12 wherein A is selected from methylene, CO, SO and SO₂; wherein X is selected from oxygen atom, methylene and >NR₁₀ with R₁₀ selected from hydrido, alkyl and benzyl; wherein each of R₁ and R₉ is independently selected from hydrido, lower alkyl, haloalkyl, cycloalkyl, alkoxycarbonyl, benzyloxycarbonyl, loweralkanoyl, alkoxyacetyl, phenyl and benzyl, and wherein the nitrogen atom to which R₁ and R₉ are attached may be combined with oxygen to form an

N-oxide; wherein each of R₂, R₄ and R₆ is independently selected from hydrido and alkyl; wherein R₃ is selected from phenylalkyl, naphthylmethyl, cyclohexylalkyl, cyclopentylalkyl, heteroarylalkyl, and

5 heteroarylcycloalkyl; wherein each of R₅ and R₈ is independently selected from



10 wherein V is selected from hydrido, alkyl, haloalkyl, benzyl and phenyl; wherein each of R₁₃ and R₁₄ is a radical independently selected from hydrido, alkyl, alkenyl, alkynyl, cycloalkyl, heteroaryl, heteroarylalkyl and heteroarylcycloalkyl; wherein R₇ is selected from substituted or unsubstituted

15 cyclohexylmethyl and benzyl, either one of which may be substituted with one or more groups selected from alkyl, hydroxy, alkoxy, halo and haloalkyl; wherein each of R₁₁ and R₁₂ is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero or one; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

20

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14. The composition of Claim 13 wherein A is selected from methylene, CO, SO and SO₂; wherein X is selected from oxygen atom, methylene and >NR₁₀ with R₁₀ selected from hydrido, alkyl and benzyl; wherein each of R₁ and R₉ is independently selected from hydrido, alkyl, alkoxyacetyl, haloalkyl, alkoxycarbonyl, benzyloxycarbonyl, and benzyl, and wherein the nitrogen

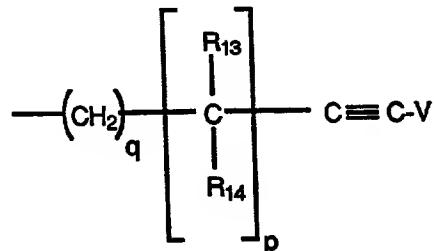
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atom to which R₁ and R₉ are attached may be combined with oxygen to form an N-oxide; wherein each of R₂, R₄ and R₆ is independently selected from hydrido and alkyl; wherein R₃ is selected from benzyl, phenethyl,

5 cyclohexylmethyl, phenpropyl, pyrrolidinyl, piperidinyl, pyrrolidinylmethyl, piperidinylmethyl, pyrazolemethyl, pyrazoleethyl, pyridylmethyl, pyridylethyl, thiazolemethyl, thiazoleethyl, imidazolemethyl, imidazoleethyl, thienylmethyl, thiencylthyl,

10 furanylmethyl, furanylethyl, oxazolemethyl, oxazoleethyl, isoxazolemethyl, isoxazoleethyl, pyridazinemethyl, pyridazineethyl, pyrazinemethyl and pyrazineethyl; wherein each of R₅ and R₈ is independently selected from

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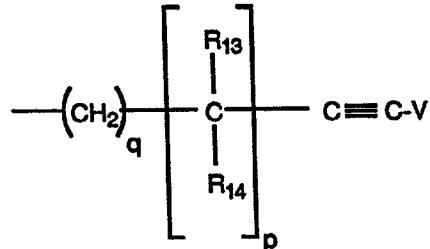


wherein V is selected from hydrido, alkyl and haloalkyl; wherein each of R₁₃ and R₁₄ is a radical independently selected from hydrido, alkyl, alkenyl, alkynyl, thiazole and thiazolemethyl; wherein R₇ is cyclohexylmethyl; wherein each of R₁₁ and R₁₂ is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero or one; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

30 15. The composition of Claim 14 wherein A is selected from CO and SO₂; wherein X is selected from oxygen atom, methylene and $\text{>} \text{NR}_{10}$ with R₁₀ selected from

hydrido and methyl; wherein each of R₁ and R₉ is independently selected from hydrido, lower alkyl, alkoxyacyl, alkoxy carbonyl, benzyloxycarbonyl, haloalkyl and benzyl, and wherein the nitrogen atom to which R₁ and R₉ are attached may be combined with oxygen to form an N-oxide; wherein R₂ is selected from hydrido, methyl, ethyl and isopropyl; wherein R₃ is selected from benzyl, phenethyl, cyclohexylmethyl, pyrrolidinyl, piperidinyl, pyrrolidinylmethyl, piperidinylmethyl, pyrazolemethyl, pyrazoleethyl, pyridylmethyl, pyridylethyl, thiazolemethyl, thiazoleethyl, imidazolemethyl, imidazoleethyl, thienylmethyl, thienylethyl, furanylmethyl, furanylethyl, oxazolemethyl, oxazoleethyl, isoxazolemethyl, isoxazoleethyl, pyridazineethyl, pyridazineethoxy, pyrazinemethyl and pyrazineethyl; wherein each of R₄ and R₆ is independently selected from hydrido and methyl; wherein each of R₅ and R₈ is independently selected from

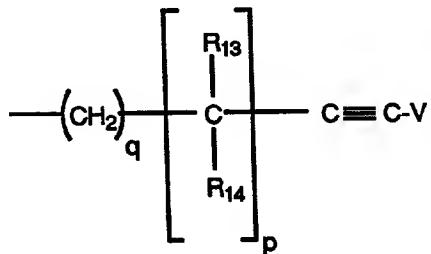
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wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R₁₃ and R₁₄ is a radical independently selected from hydrido, alkyl and alkynyl; wherein R₇ is cyclohexylmethyl; wherein each of R₁₁ and R₁₂ is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

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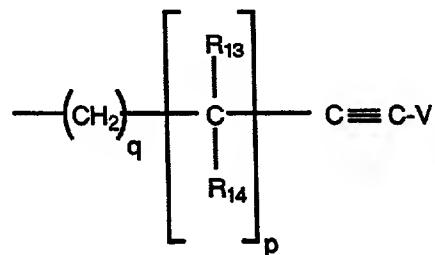
16. The composition of Claim 15 wherein A is selected from CO and SO₂; wherein X is selected from oxygen atom and methylene; wherein each of R₁ and R₉ is independently selected from hydrido, methyl, ethyl, n-propyl, isopropyl, benzyl, b, b, b-trifluoroethyl, t-butyloxycarbonyl and methoxymethylcarbonyl, and wherein the nitrogen atom to which R₁ and R₉ are attached may be combined with oxygen to form an N-oxide; wherein R₂ is selected from hydrido, methyl, ethyl and isopropyl; wherein R₃ is selected from benzyl, cyclohexylmethyl, phenethyl, pyrazolemethyl, pyrazoleethyl, pyridylmethyl, pyridylethyl, thiazolemethyl, thiazoleethyl, imidazolemethyl, imidazoleethyl, thiethylmethyl, thienelethyl, furanylmethyl, furanylethyl, oxazolemethyl, oxazoleethyl, isoxazolemethyl, isoxazoleethyl, pyridazinemethyl, pyridazineethyl, pyrazinemethyl and pyrazineethyl; wherein each of R₅ and R₈ is independently selected from



wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R₁₃ and R₁₄ is a radical independently selected from hydrido, methyl, ethyl, propyl and ethynyl; wherein R₇ is cyclohexylmethyl; wherein each of R₄ and R₆ is independently selected from hydrido and methyl; wherein each of R₁₁ and R₁₂ is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero; wherein n is a number selected from zero through five; wherein p is a number selected from zero through

five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

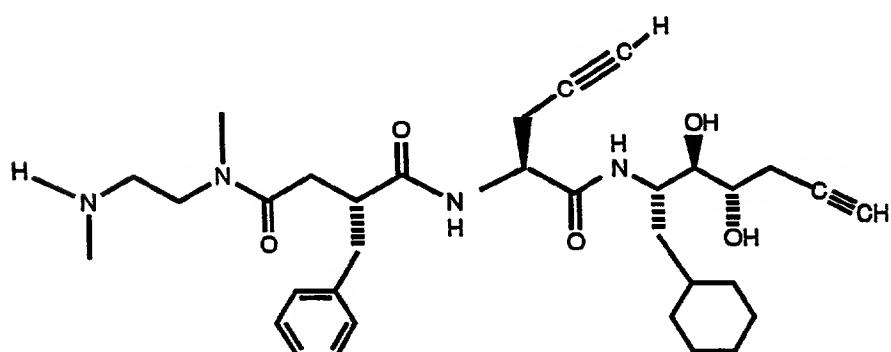
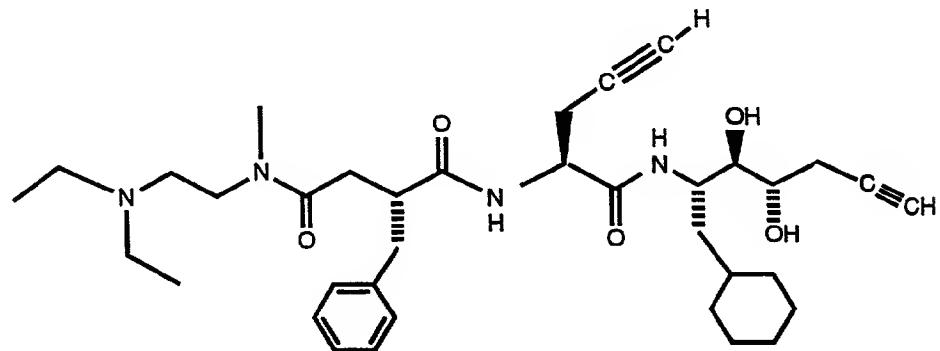
5 17. The composition of Claim 16 wherein A is selected from CO and SO₂; wherein X is selected from oxygen atom and methylene; wherein each of R₁ and R₉ is a group independently selected from hydrido, methyl, ethyl, n-propyl, isopropyl, benzyl, b, b,
10 b-trifluoroethyl, t-butyloxycarbonyl and methoxymethylcarbonyl, and wherein the nitrogen atom to which R₁ and R₉ are attached may be combined with oxygen to form an N-oxide; wherein R₂ is selected from hydrido, methyl, ethyl and isopropyl; wherein R₃ is selected from benzyl, cyclohexylmethyl, phenethyl, imidazolemethyl, pyridylmethyl and 2-pyridylethyl; wherein each of R₅ and R₈ is independently selected from
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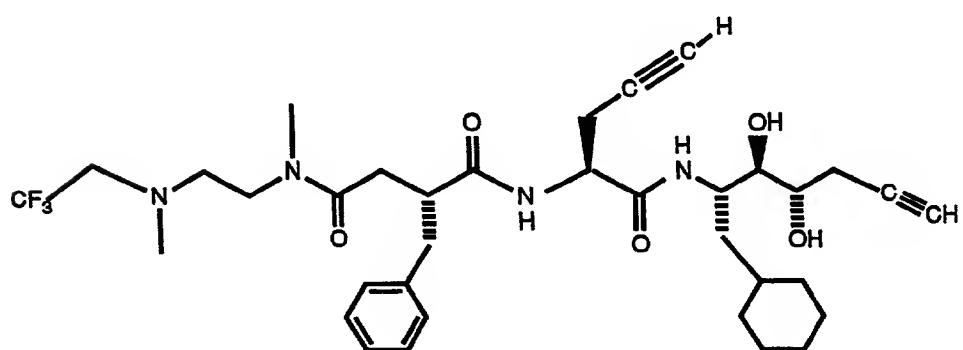
20 wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R₁₃ and R₁₄ is a radical independently selected from hydrido, methyl and ethynyl; wherein R₇ is cyclohexylmethyl; wherein each of R₄ and R₆ is independently selected from hydrido and methyl; wherein each of R₁₁ and R₁₂ is independently selected from hydrido, alkyl and phenyl; wherein m is 25 zero; wherein n is a number selected from zero through three; wherein p is a number selected from one through three; and wherein q is zero or one; or a pharmaceutically-acceptable salt thereof.

18. The composition of Claim 17 wherein said renin-inhibiting compound is selected from compounds, their tautomers, and the pharmaceutically-acceptable esters and salts thereof, of the group consisting of

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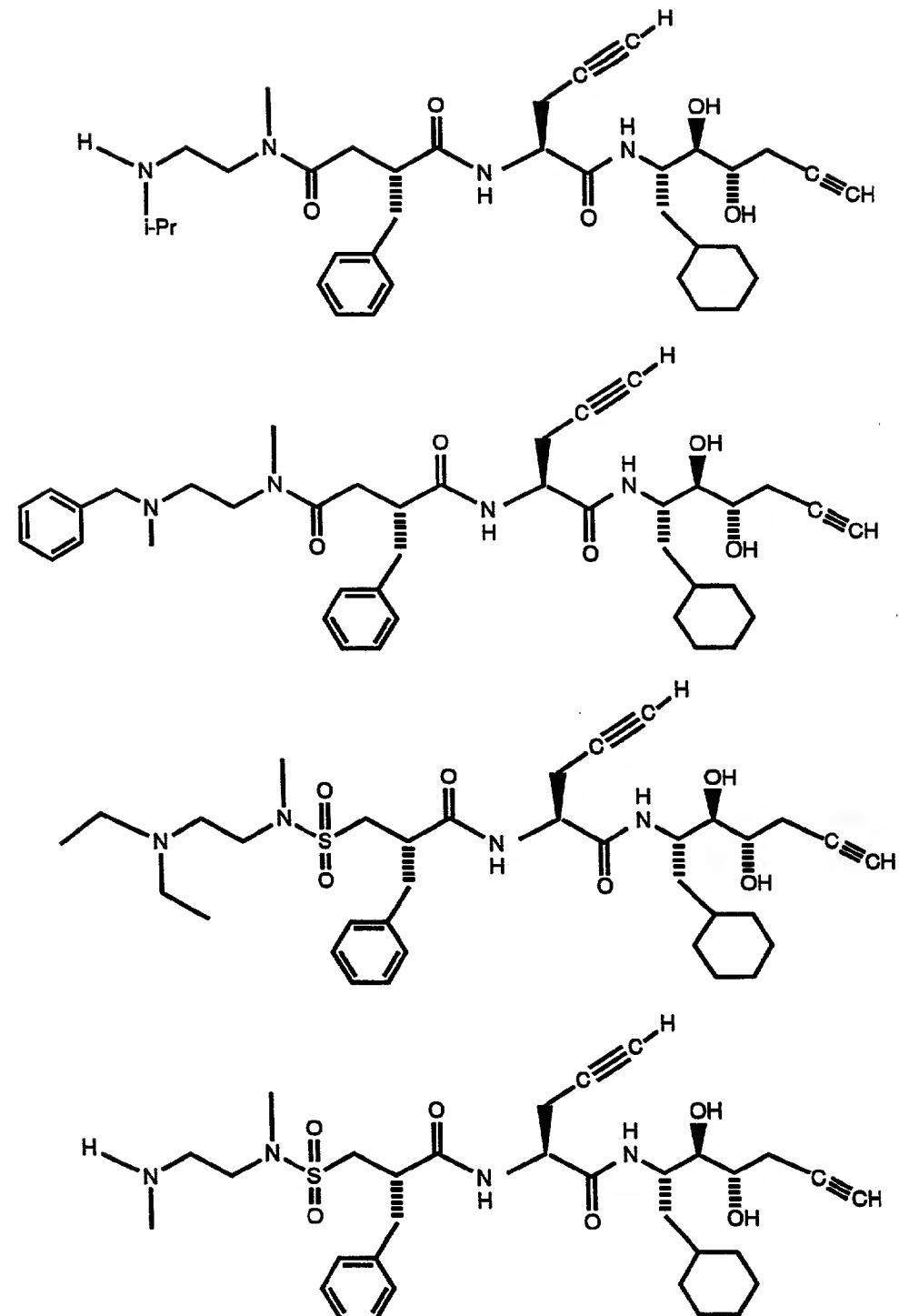


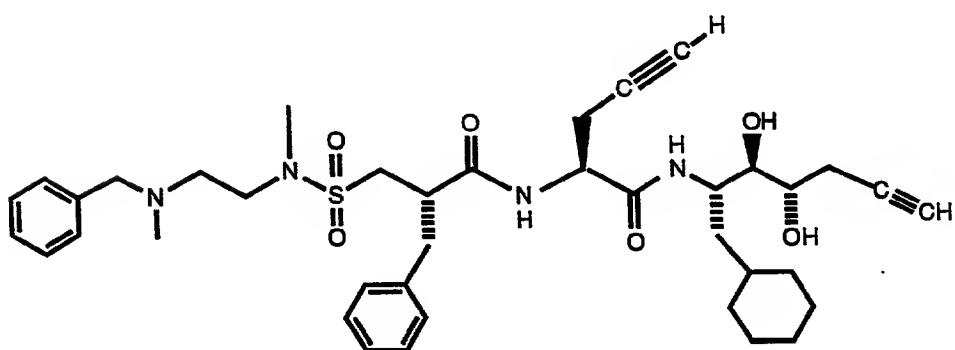
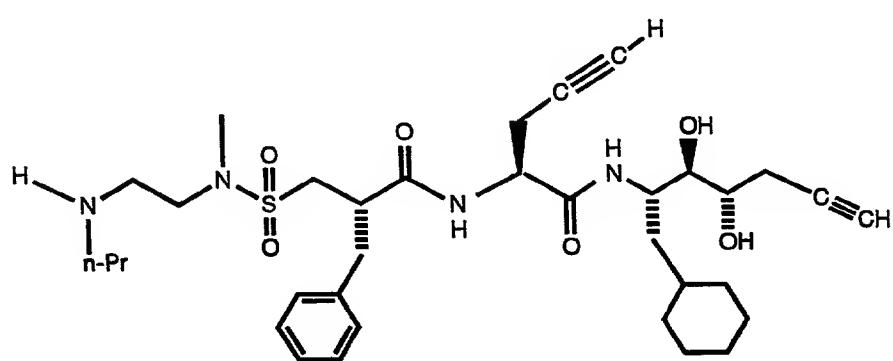
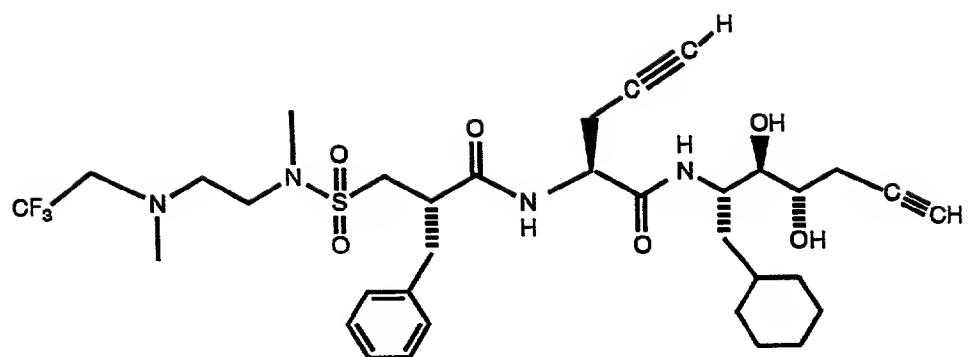
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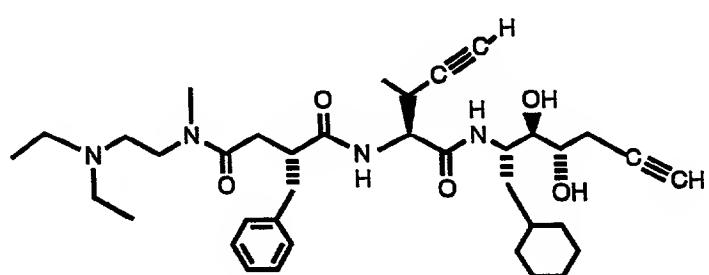
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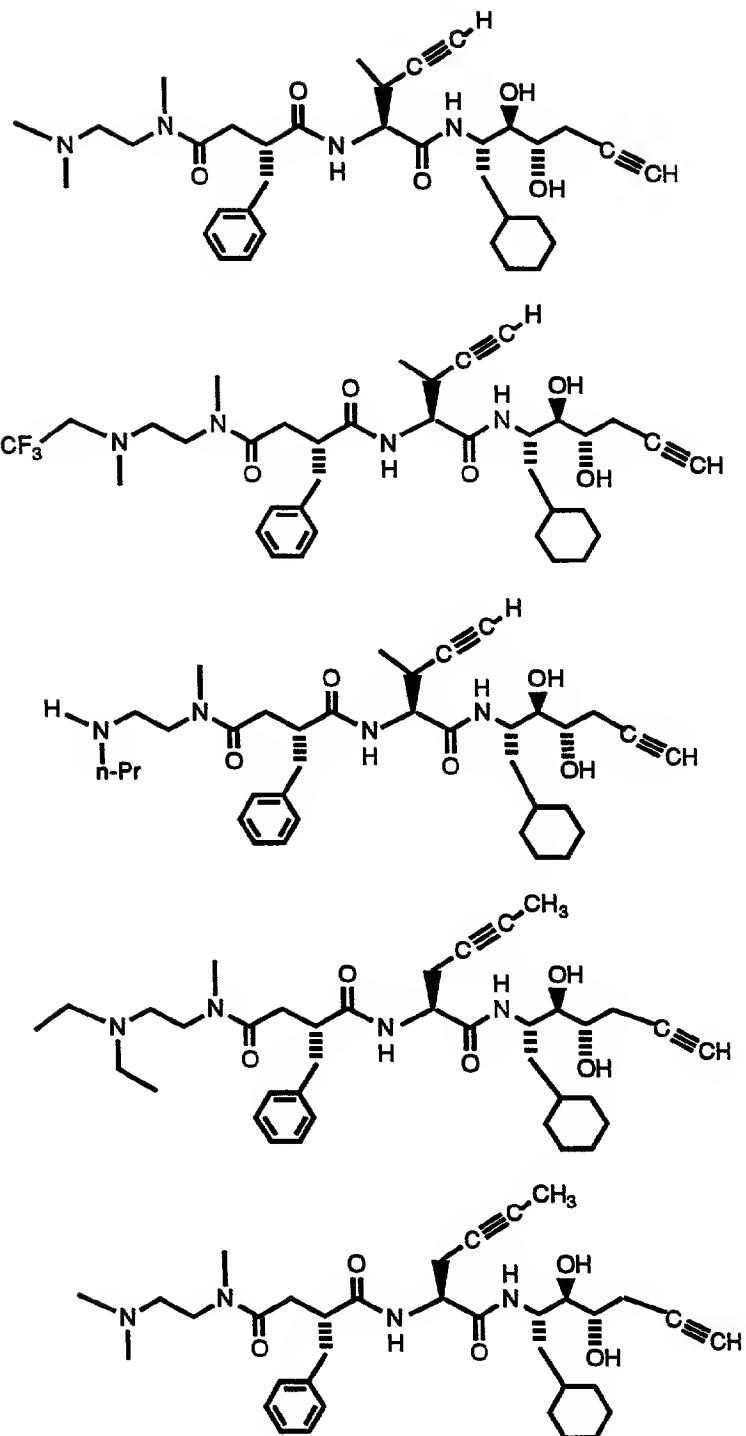
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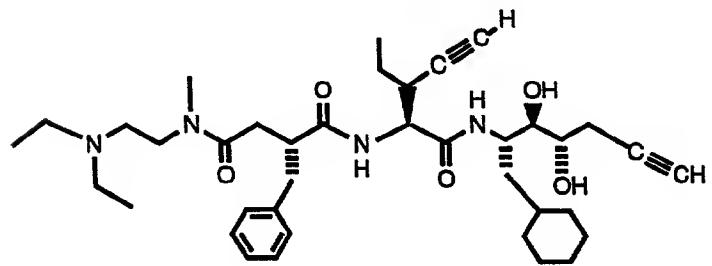
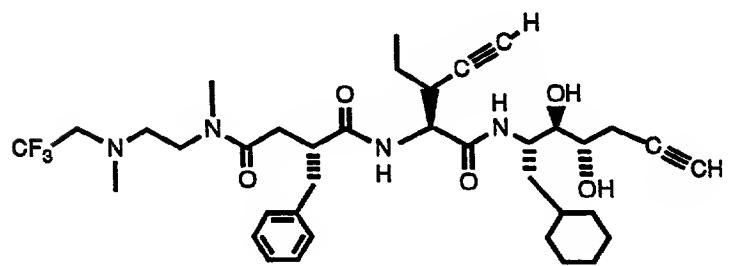
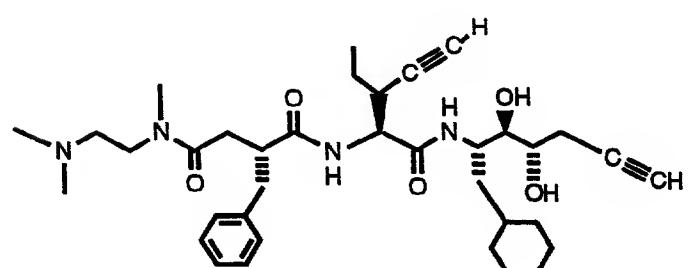
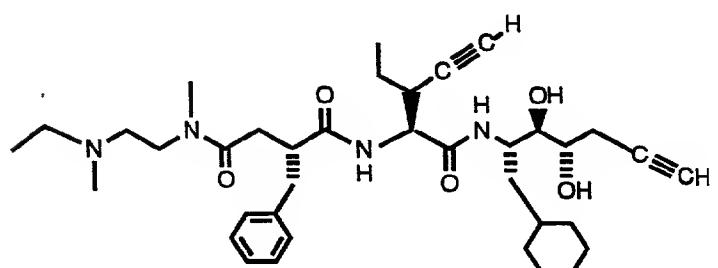
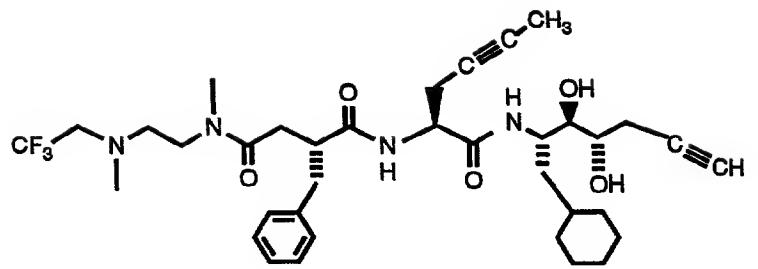




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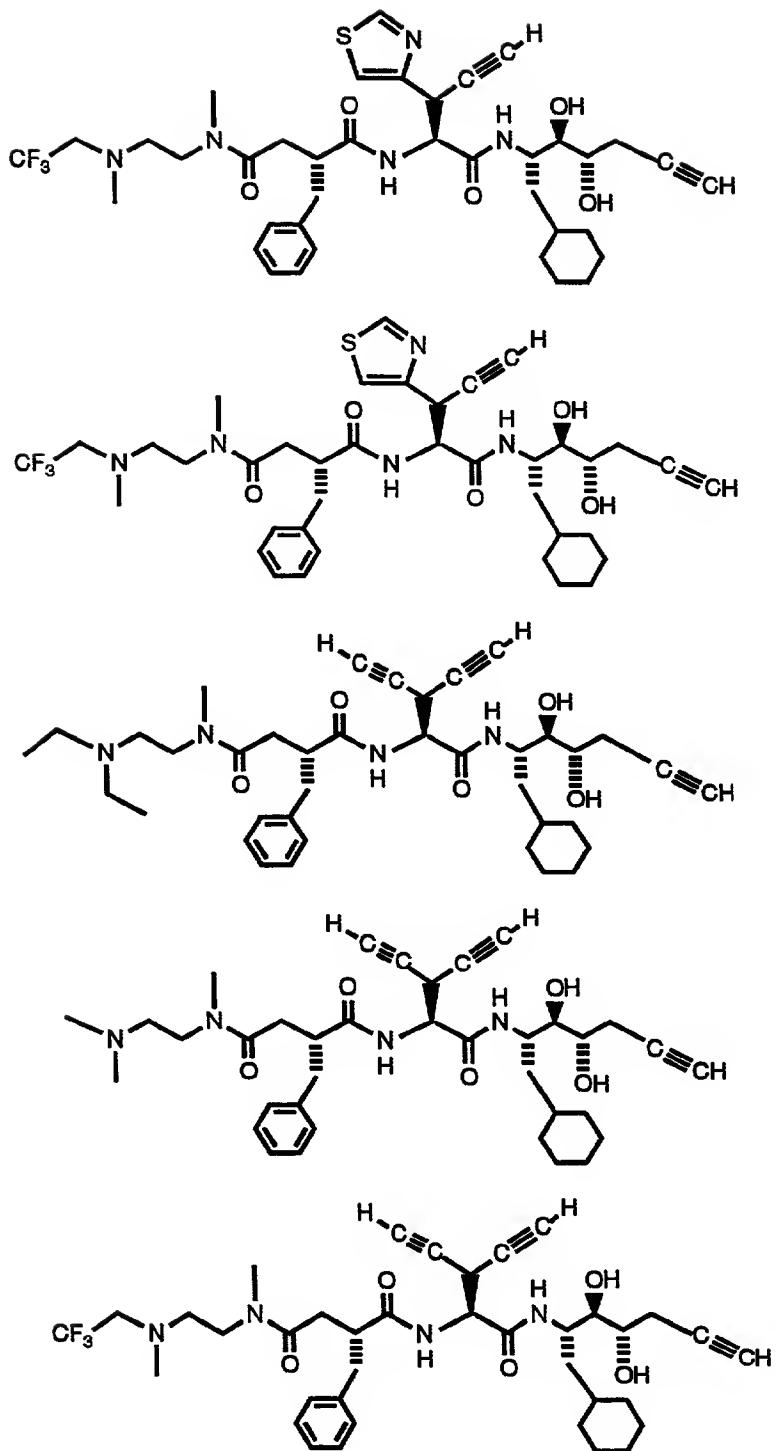
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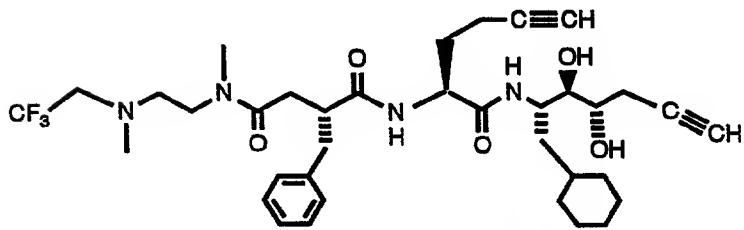
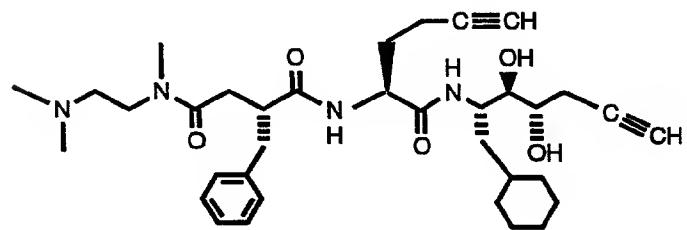
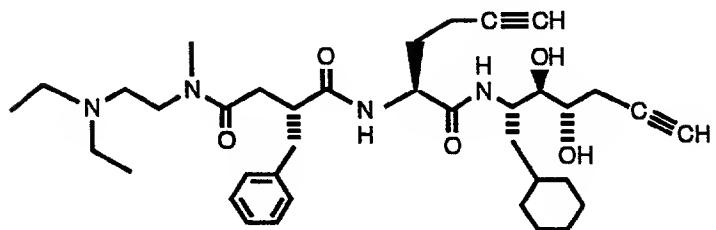
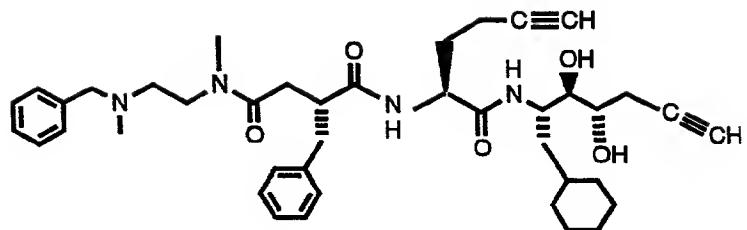
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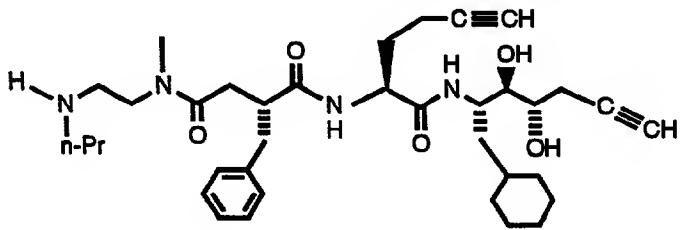
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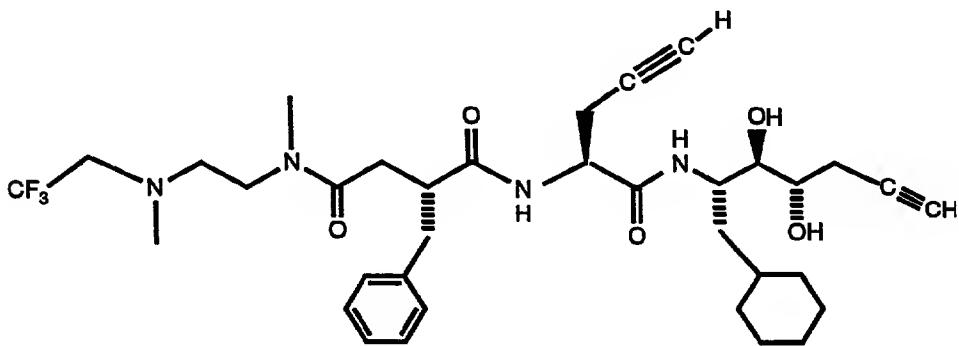
and



19. The composition of Claim 17 wherein said renin-inhibiting compound is N1-[1R*-[[1S,1R*-(cyclohexylmethyl)-2S*,3R*-dihydroxyhexynyl]amino]carbonyl]-3-butynyl]-N4-[2-
5 (dimethylamino)ethyl]-N4-methyl-2S*-(phenylmethyl)butanediamide or a pharmaceutically-acceptable salt thereof.

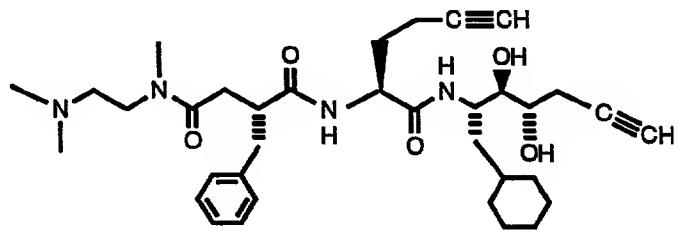
10 20. The composition of Claim 17 wherein said renin-inhibiting compound is [1R*-[[1R*-[[1S,1R*-(cyclohexylmethyl)-2S*,3R*-dihydroxyhexynyl]amino]carbonyl]-3-butynyl]amino]carbonyl]-2-phenylethyl)[2-(dimethylamino)ethyl]methylcarbamate or a
15 pharmaceutically-acceptable salt thereof.

21. The composition of Claim 17 wherein said renin-inhibiting compound is



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22. The composition of Claim 17 wherein said renin-inhibiting compound is

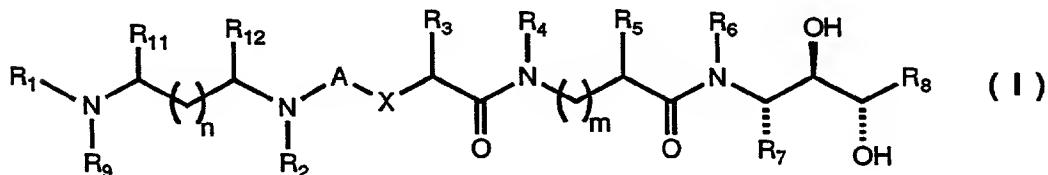


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or a pharmaceutically-acceptable salt thereof.

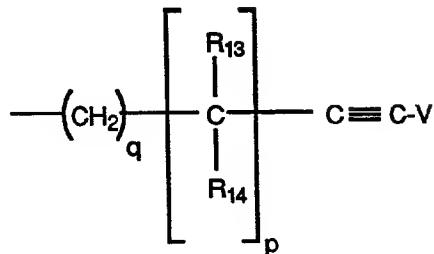
23. A therapeutic method for treating a circulatory disorder or a circulatory-related disorder, said method comprising administering to a subject susceptible to or
 5 afflicted with such disorder a therapeutically-effective amount of an active compound

of Formula I:



wherein A is selected from methylene, CO, SO and SO₂;
 wherein X is selected from oxygen atom, methylene and
 >NR₁₀ with R₁₀ selected from hydrido, alkyl and benzyl;
 wherein each of R₁ and R₉ is a group independently selected from hydrido, alkyl, cycloalkyl, alkoxyacetyl, haloalkyl, alkoxycarbonyl, benzyloxycarbonyl, loweralkanoyl, haloalkylacyl, phenyl, benzyl, naphthyl, and naphthylmethyl, any one of which groups having a substitutable position may be optionally substituted
 20 with one or more radicals selected from alkyl, alkoxy, alkenyl, alkynyl, halo, haloalkyl, cyano and phenyl, and wherein the nitrogen atom to which R₁ and R₉ are attached may be combined with oxygen to form an N-oxide;
 wherein R₂ is selected from hydrido, alkyl,
 25 dialkylaminoalkyl, alkylacylaminoalkyl, benzyl and cycloalkyl; wherein R₃ is selected from alkyl, cycloalkylalkyl, acylaminoalkyl, phenylalkyl, naphthylmethyl, aryl, heterocyclicalkyl and heterocycliccycloalkyl, wherein the cyclic portion of
 30 any of said phenylalkyl, naphthylmethyl, aryl, heterocyclicalkyl and heterocycliccycloalkyl groups may be substituted by one or more radicals selected from halo, hydroxy, alkoxy and alkyl; wherein each of R₄ and R₆ is independently selected from hydrido, alkyl, benzyl

and cycloalkyl; wherein each of R₅ and R₈ is independently selected from



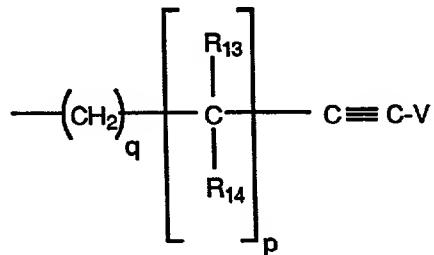
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wherein V is selected from hydrido, alkyl, cycloalkyl, haloalkyl, benzyl and phenyl; wherein each of R₁₃ and R₁₄ is a radical independently selected from hydrido, alkyl, alkenyl, alkynyl, cycloalkyl, phenyl, heterocyclic, heterocyclicalkyl and heterocycliccycloalkyl; wherein R₇ is selected from substituted or unsubstituted alkyl, cycloalkyl, phenyl, cycloalkylalkyl and phenylalkyl, any one of which may be substituted with one or more groups selected from alkyl, hydroxy, alkoxy, halo, haloalkyl, alkenyl, alkynyl and cyano; wherein each of R₁₁ and R₁₂ is independently selected from hydrido, alkyl, haloalkyl, dialkylamino and phenyl; and wherein m is zero or one; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

24. The method of Claim 23 wherein A is selected from methylene, CO, SO and SO₂; wherein X is selected from oxygen atom, methylene and >NR₁₀ with R₁₀ selected from hydrido, alkyl and benzyl; wherein each of R₁ and R₉ is independently selected from hydrido, lower alkyl, haloalkyl, cycloalkyl, alkoxycarbonyl, benzyloxycarbonyl, loweralkanoyl, alkoxyacetyl, phenyl and benzyl, and wherein the nitrogen atom to which R₁ and R₉ are attached may be combined with oxygen to form an

N-oxide; wherein each of R₂, R₄ and R₆ is independently selected from hydrido and alkyl; wherein R₃ is selected from phenylalkyl, naphthylmethyl, cyclohexylalkyl, cyclopentylalkyl, heteroarylalkyl,

5 heteroarylcycloalkyl; wherein each of R₅ and R₈ is independently selected from



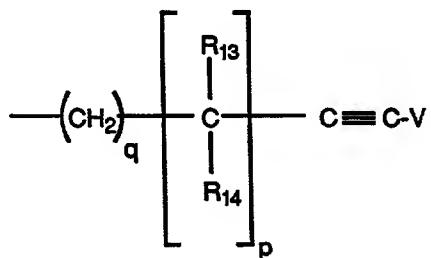
10 wherein V is selected from hydrido, alkyl, haloalkyl, benzyl and phenyl; wherein each of R₁₃ and R₁₄ is a radical independently selected from hydrido, alkyl, alkenyl, alkynyl, cycloalkyl, heteroaryl, heteroarylalkyl and heteroarylcycloalkyl; wherein R₇ is selected from substituted or unsubstituted

15 cyclohexylmethyl and benzyl, either one of which may be substituted with one or more groups selected from alkyl, hydroxy, alkoxy, halo and haloalkyl; wherein each of R₁₁ and R₁₂ is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero or one; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

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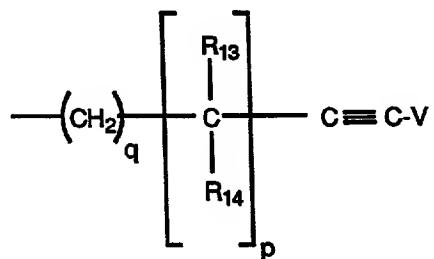
25. The method of Claim 24 wherein A is selected from methylene, CO, SO and SO₂; wherein X is selected from oxygen atom, methylene and >NR₁₀ with R₁₀ selected from hyrido, alkyl and benzyl; wherein each of 5 R₁ and R₉ is independently selected from hydrido, alkyl, alkoxyacyl, haloalkyl, alkoxycarbonyl, benzyloxycarbonyl and benzyl, and wherein the nitrogen atom to which R₁ and R₉ are attached may be combined with oxygen to form an N-oxide; wherein each of R₂, R₄ and R₆ is 10 independently selected from hydrido and alkyl; wherein R₃ is selected from benzyl, phenethyl, cyclohexylmethyl, phenpropyl, pyrrolidinyl, piperidinyl, pyrrolidinylmethyl, piperidinylmethyl, pyrazolemethyl, pyrazoleethyl, pyridylmethyl, pyridylethyl, 15 thiazolemethyl, thiazoleethyl, imidazolemethyl, imidazoleethyl, thienylmethyl, thiencyethyl, furanylmethyl, furanylethyl, oxazolemethyl, oxazoleethyl, isoxazolemethyl, isoxazoleethyl, pyridazinemethyl, pyridazineethyl, pyrazinemethyl and pyrazineethyl; wherein each of R₅ and R₈ is 20 independently selected from



25. wherein V is selected from hydrido, alkyl and haloalkyl; wherein each of R₁₃ and R₁₄ is a radical independently selected from hydrido, alkyl, alkenyl, alkynyl, thiazole and thiazolemethyl; wherein R₇ is cyclohexylmethyl; wherein each of R₁₁ and R₁₂ is independently selected 30 from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero or one; wherein n is a number selected from zero through five; wherein p is a number selected from zero

through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

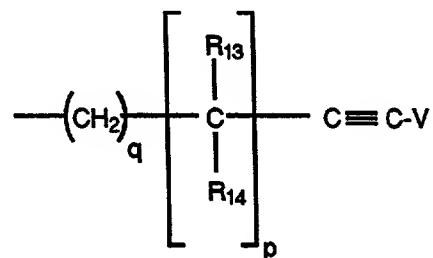
5 26. The method of Claim 25 wherein A is
selected from CO and SO₂; wherein X is selected from
oxygen atom, methylene and >NR₁₀ with R₁₀ selected from
hydrido and methyl; wherein each of R₁ and R₉ is
independently selected from hydrido, lower alkyl,
10 alkoxyacyl, alkoxycarbonyl, benzyloxycarbonyl, haloalkyl
and benzyl, and wherein the nitrogen atom to which R₁
and R₉ are attached may be combined with oxygen to form
an N-oxide; wherein R₂ is selected from hydrido, methyl,
ethyl and isopropyl; wherein R₃ is selected from benzyl,
15 phenethyl, cyclohexylmethyl, pyrrolidinyl, piperidinyl,
pyrrolidinylmethyl, piperidinylmethyl, pyrazolemethyl,
pyrazoleethyl, pyridylmethyl, pyridylethyl,
thiazolemethyl, thiazoleethyl, imidazolemethyl,
imidazoleethyl, thiencylmethyl, thiencylethyl,
20 furanymethyl, furanylethyl, oxazolemethyl,
oxazoleethyl, isoxazolemethyl, isoxazoleethyl,
pyridazinemethyl, pyridazineethyl, pyrazinemethyl and
pyrazineethyl; wherein each of R₄ and R₆ is
independently selected from hydrido and methyl; wherein
25 each of R₅ and R₈ is independently selected from



wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R₁₃ and R₁₄ is a radical independently selected from hydrido, alkyl and alkynyl; wherein R₇ is cyclohexylmethyl; wherein each of

R₁₁ and R₁₂ is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; 5 and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

27. The method of Claim 26 wherein A is selected from CO and SO₂; wherein X is selected from oxygen atom and methylene; wherein each of R₁ and R₉ is independently selected from hydrido, methyl, ethyl, n-propyl, isopropyl, benzyl, b, b, b-trifluoroethyl, t-butyloxycarbonyl and methoxymethylcarbonyl, and wherein the nitrogen atom to which R₁ and R₉ are attached may be combined with oxygen to form an N-oxide; 10 wherein R₂ is selected from hydrido, methyl, ethyl and isopropyl; wherein R₃ is selected from benzyl, cyclohexylmethyl, phenethyl, pyrazolemethyl, pyrazoleethyl, pyridylmethyl, pyridylethyl, 15 thiazolemethyl, thiazoleethyl, imidazolemethyl, imidazoleethyl, thienylmethyl, thiénylethyl, furanylmethyl, furanylethyl, oxazolemethyl, oxazoleethyl, isoxazolemethyl, isoxazoleethyl, pyridazinemethyl, pyridazineethyl, pyrazinemethyl and 20 pyrazineethyl; wherein each of R₅ and R₈ is independently selected from 25

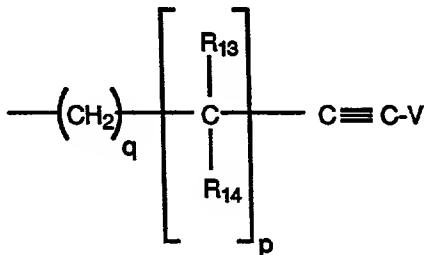


30 wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R₁₃ and R₁₄ is a radical independently selected from hydrido, methyl, ethyl, propyl and ethynyl; wherein R₇ is

cyclohexylmethyl; wherein each of R₄ and R₆ is independently selected from hydrido and methyl; wherein each of R₁₁ and R₁₂ is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

10

28. The method of Claim 27 wherein A is selected from CO and SO₂; wherein X is selected from oxygen atom and methylene; wherein each of R₁ and R₉ is a group independently selected from hydrido, methyl, ethyl, n-propyl, isopropyl, benzyl, b, b, b-trifluoroethyl, t-butyloxycarbonyl and methoxymethylcarbonyl, and wherein the nitrogen atom to which R₁ and R₉ are attached may be combined with oxygen to form an N-oxide; wherein R₂ is selected from hydrido, methyl, ethyl and isopropyl; wherein R₃ is selected from benzyl, cyclohexylmethyl, phenethyl, imidazolemethyl, pyridylmethyl and 2-pyridylethyl; wherein each of R₅ and R₈ is independently selected from



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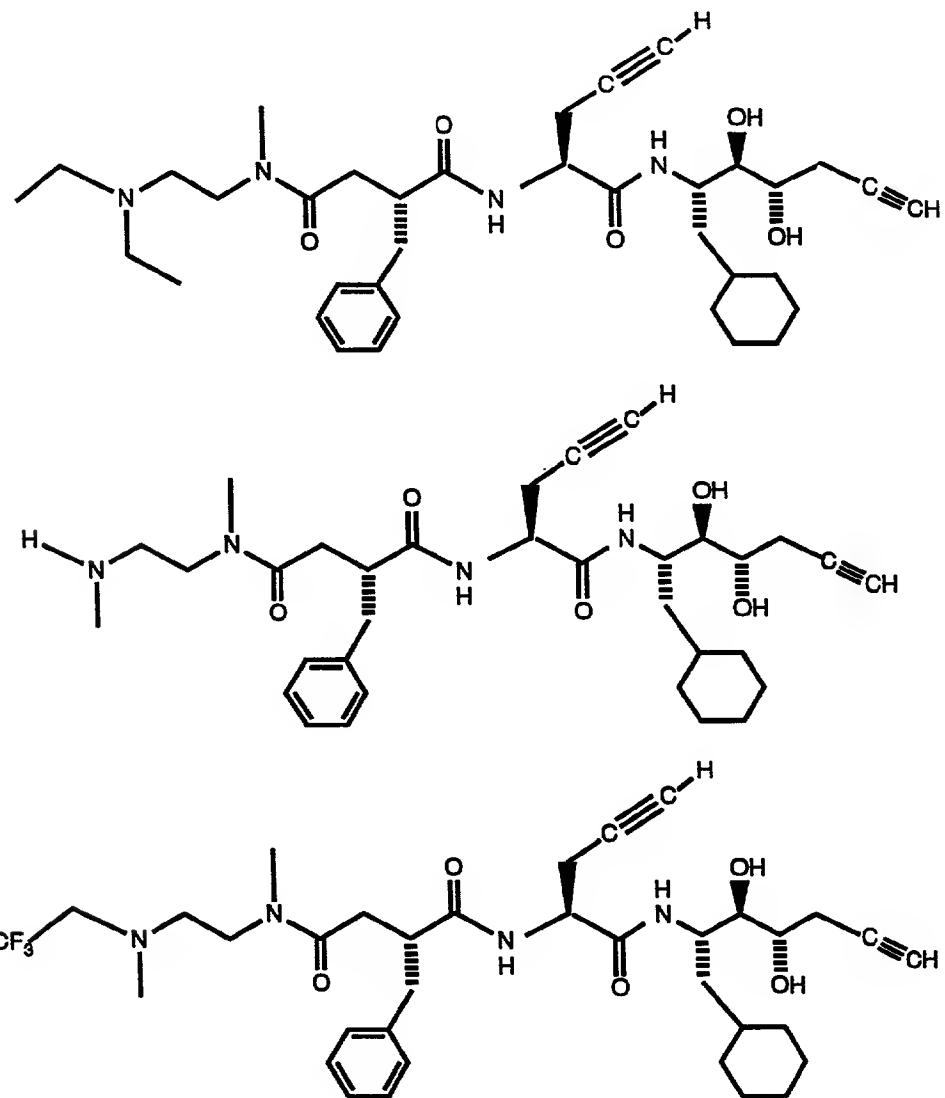
wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R₁₃ and R₁₄ is a radical independently selected from hydrido, methyl and ethynyl; wherein R₇ is cyclohexylmethyl; wherein each of R₄ and R₆ is independently selected from hydrido and methyl; wherein each of R₁₁ and R₁₂ is independently selected from hydrido, alkyl and phenyl; wherein m is

zero; wherein n is a number selected from zero through three; wherein p is a number selected from one through three; and wherein q is zero or one; or a pharmaceutically-acceptable salt thereof.

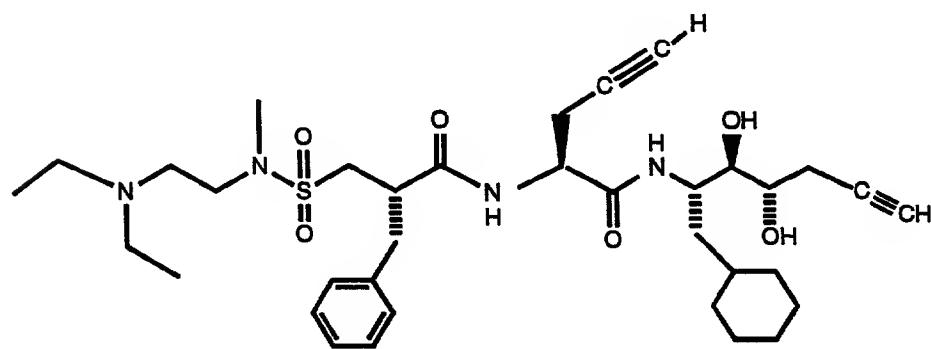
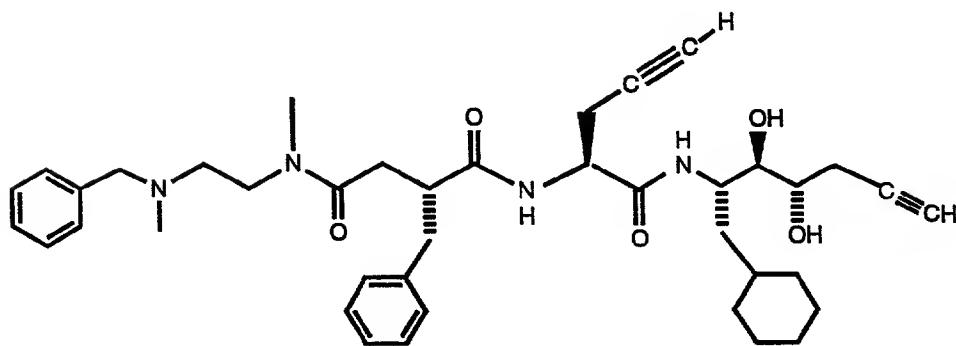
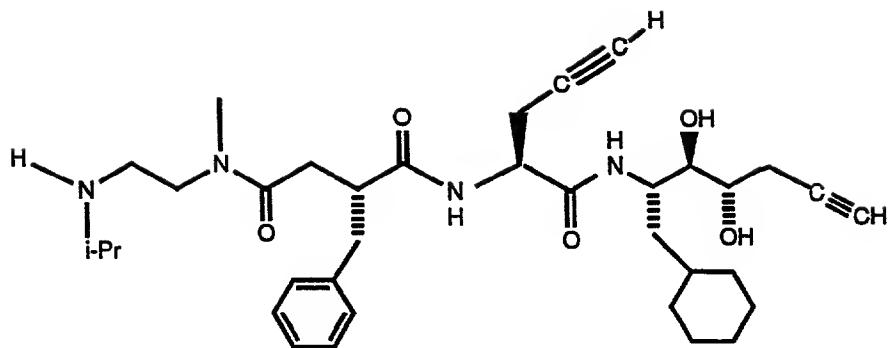
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29. The method of Claim 28 wherein said compound is selected from compounds, their tautomers, and the pharmaceutically-acceptable esters and salts thereof, of the group consisting of

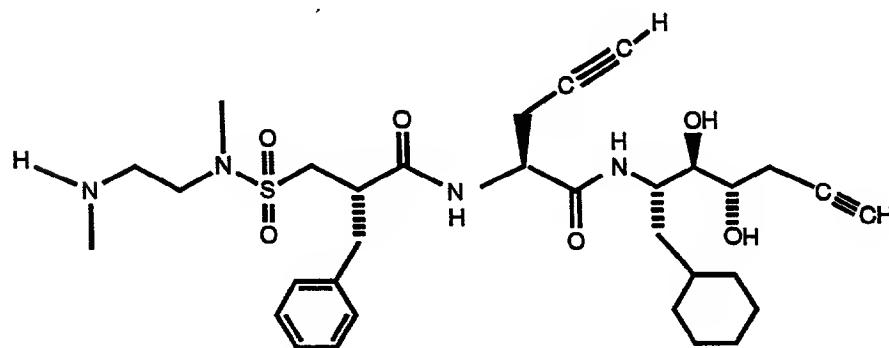
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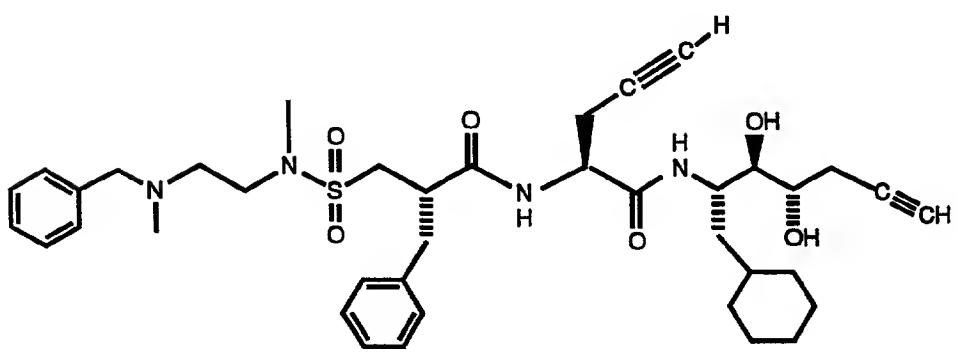
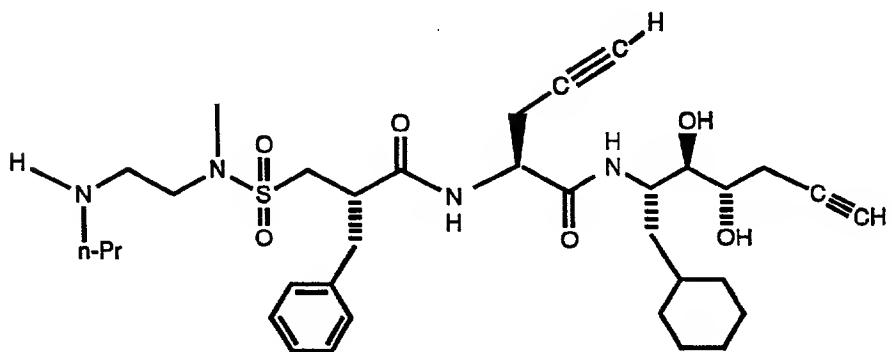
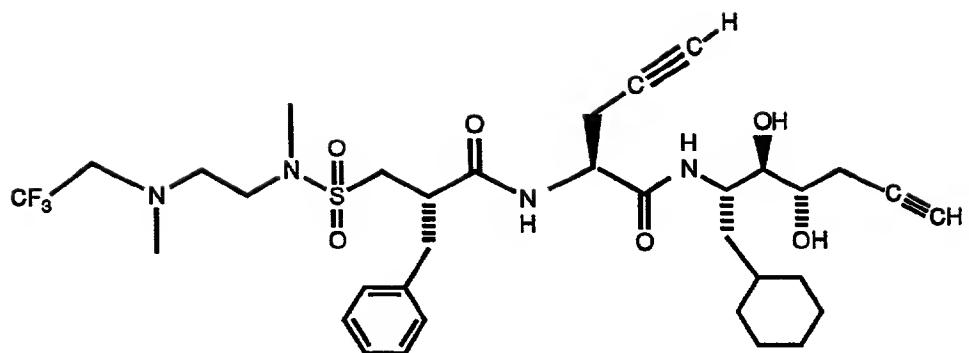
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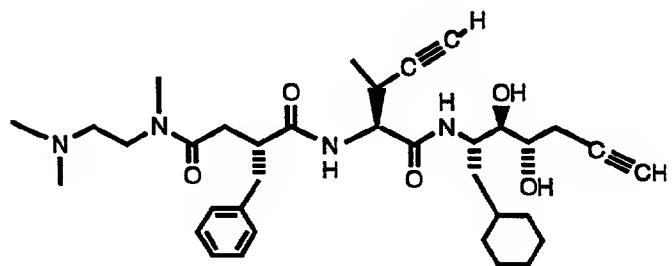
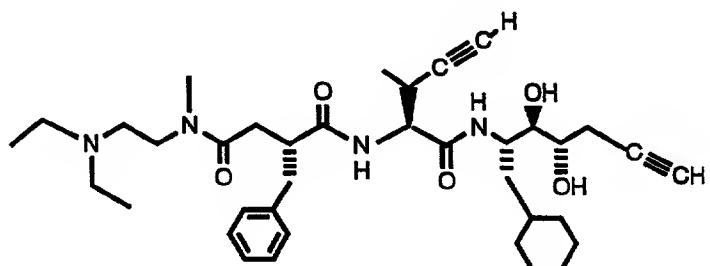
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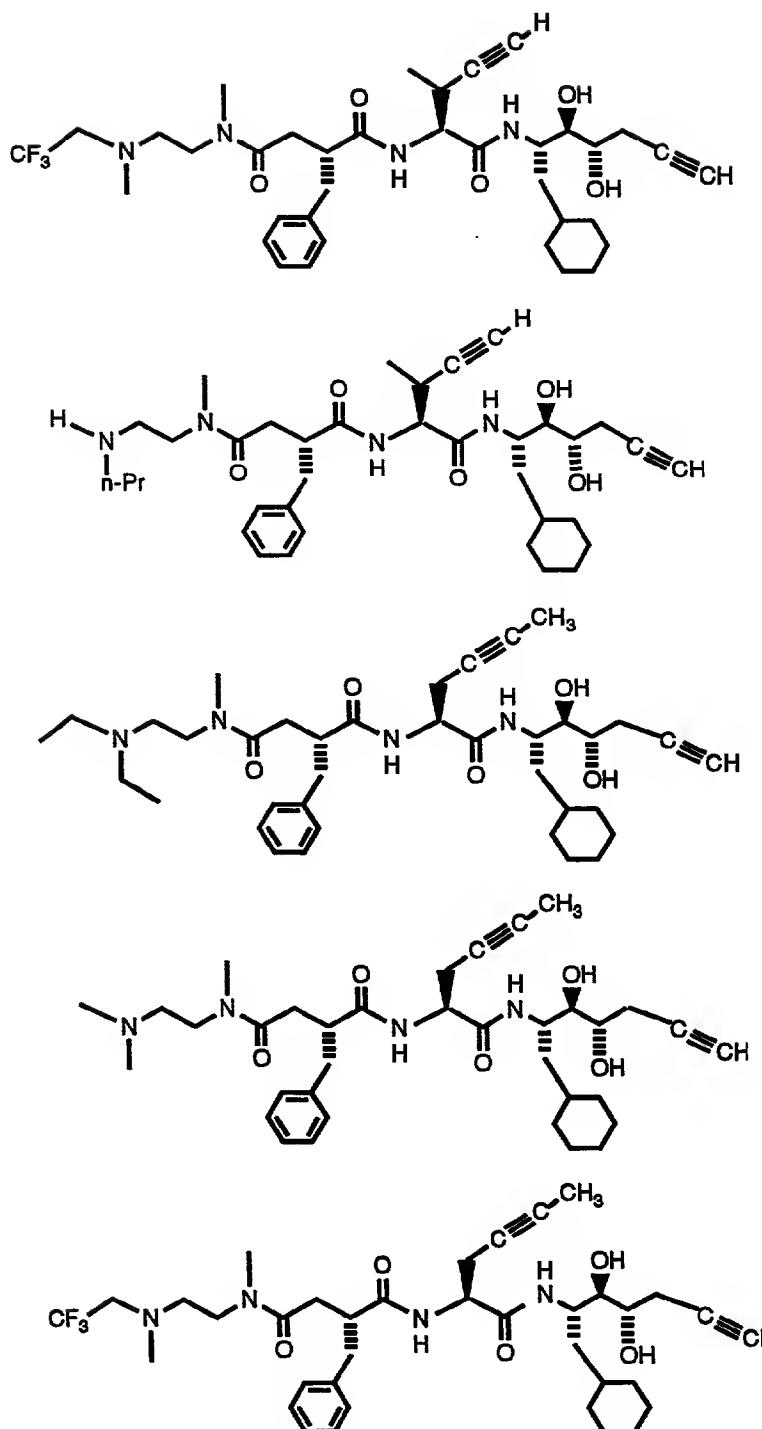


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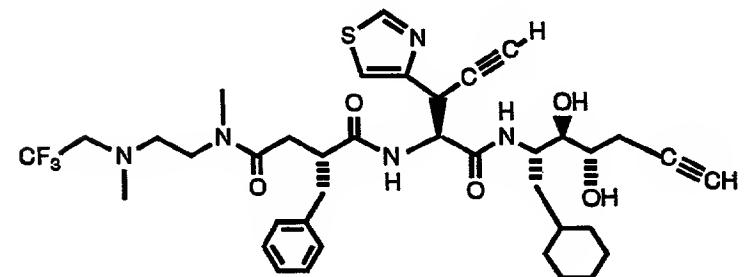
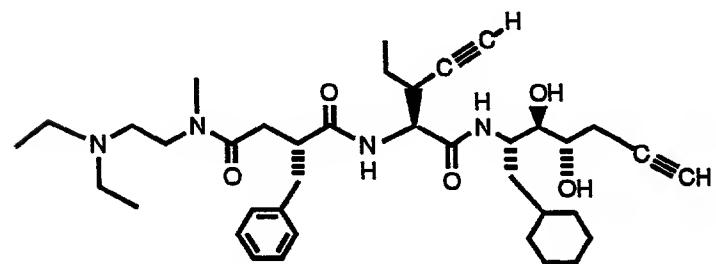
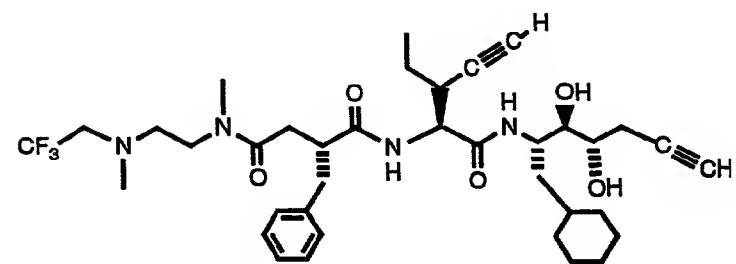
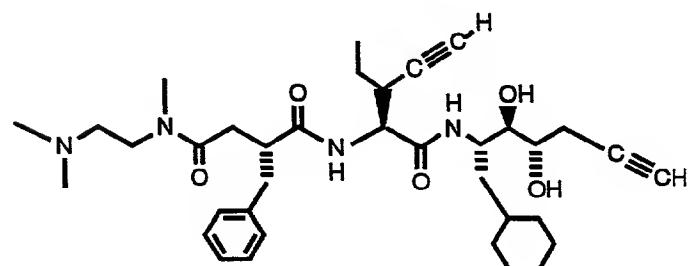
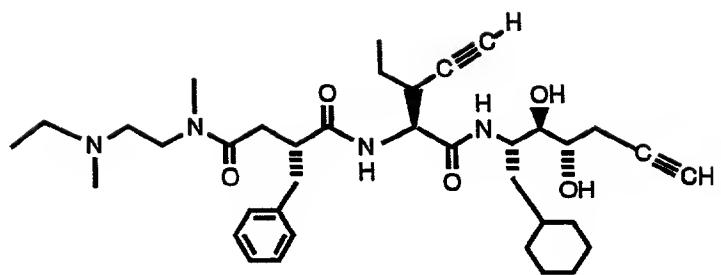


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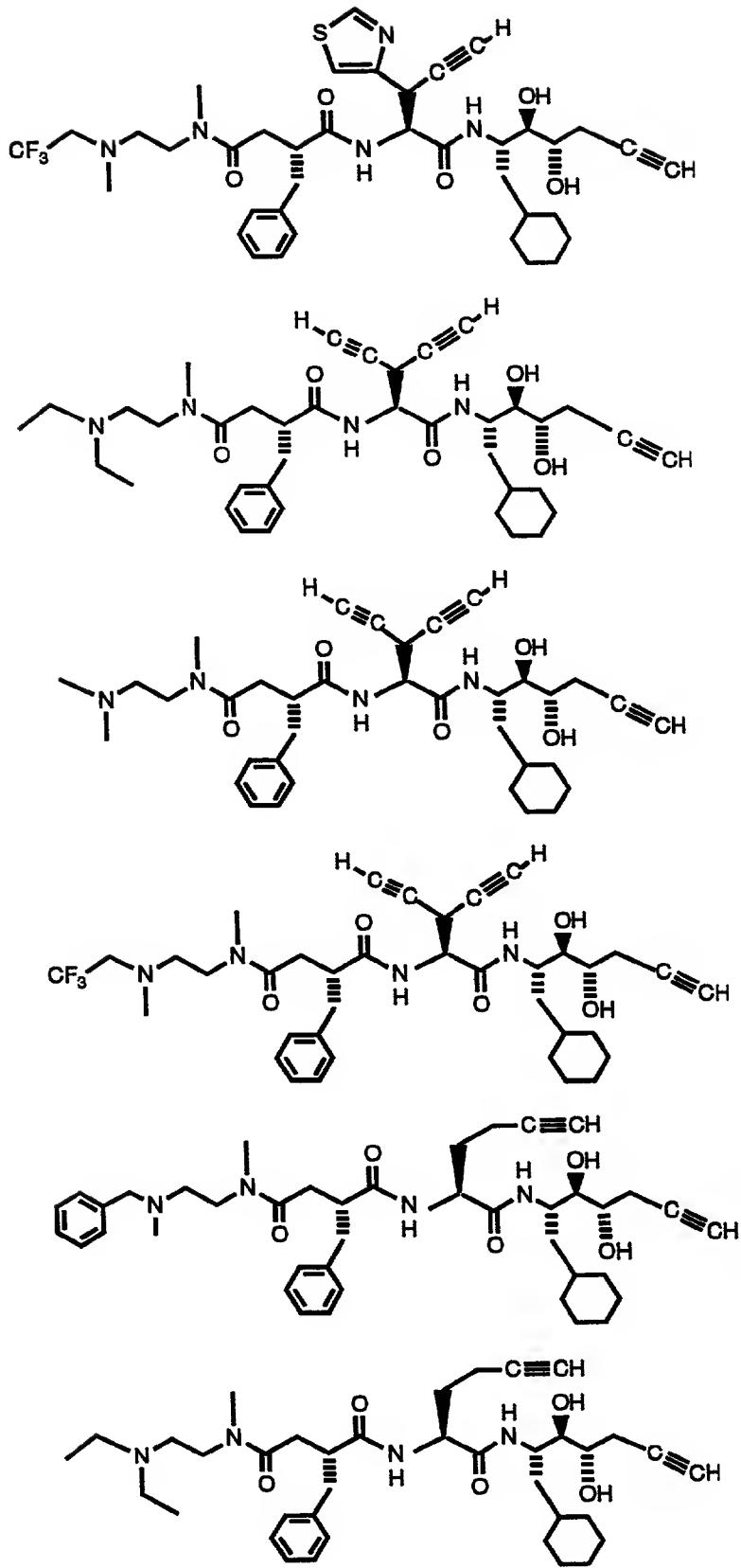


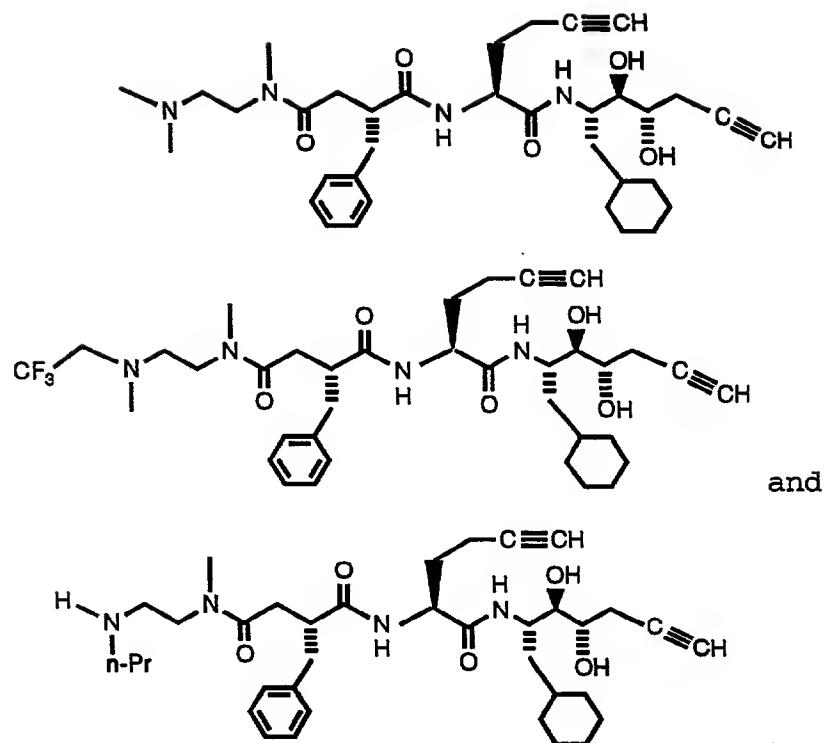
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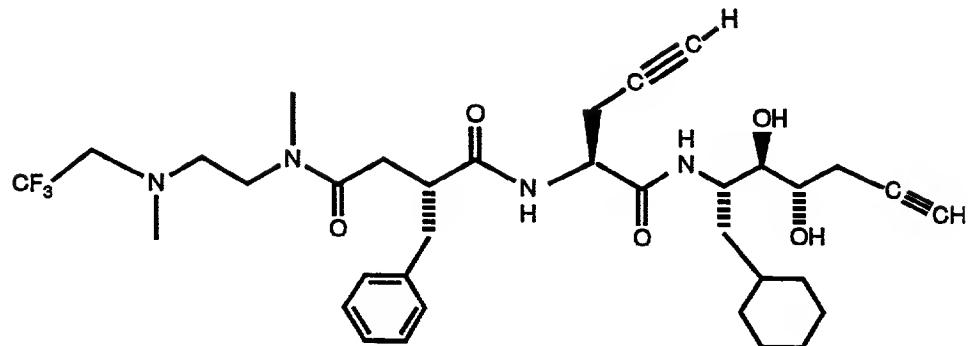




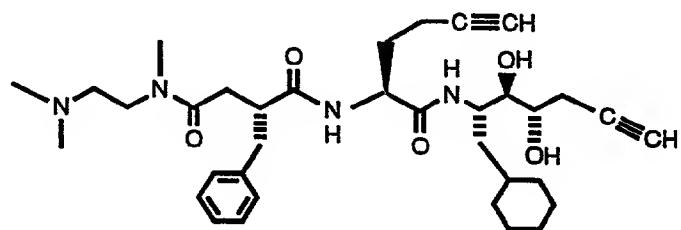
30. The method of Claim 28 wherein said compound is N1-[1R*-[[1S,1R*-(cyclohexylmethyl)-2S*,3R*-dihydroxy-hexynyl]amino]carbonyl]-3-butynyl]-N4-[2-(dimethylamino)ethyl]-N4-methyl-2S*(phenylmethyl)butanediamide or a pharmaceutically-acceptable salt thereof.

31. The method of Claim 28 wherein said compound is [1R*-[[1R*-[[1S,1R*-(cyclohexylmethyl)-2S*,3R*-dihydroxy-hexynyl]amino]carbonyl]-3-butynyl]amino]carbonyl]-2-phenylethyl]-N4-[2-(dimethylamino)ethyl]-N4-methyl-2S*(phenylmethyl)butanediamide or a pharmaceutically-acceptable salt thereof.

32. The method of Claim 28 wherein said compound is



5 33. The method of Claim 28 wherein said compound is



10 or a pharmaceutically-acceptable salt thereof.

34. The method of Claim 23 wherein said circulatory disorder is a cardiovascular disorder.

15 35. The method of Claim 34 wherein said cardiovascular disorder is hypertension.

20 36. The method of Claim 23 wherein said circulatory-related disorder is glaucoma.